

10-5-00

A

Patent  
030591.0010.CIP1

To: Box Patent Application  
Commissioner for Patents  
Washington, D.C. 20231

JB913 U.S. PTO  
09/679664  
10/03/00

**CONTINUATION-IN-PART APPLICATION**

**TRANSMITTAL**

Sir:

Transmitted herewith for filing is a **Continuation-in-Part** of International Application No. PCT/US99/07333 which claims the benefit of U.S. Application No. 60/080,671, Filed April 3, 1998.

**Inventor(s): Thomas Stormann, Lance G. Hammerland, Laura L. Storjohann, James G. Busby, James E. Garrett, Rachel T. Simin**

**Title: G-PROTEIN FUSION RECEPTORS AND CHIMERIC GABA<sub>B</sub> RECEPTORS**

**I. PAPERS ENCLOSED HEREWITH FOR FILING UNDER 37 CFR § 1.53(b):**

- 33 Page(s) of Written Description
- 7 Page(s) Claims
- 1 Page(s) Abstract
- 102 Other: Sequence Listing
- 116 Sheets of Drawings \_\_\_\_\_ Informal X Formal

**II. ADDITIONAL PAPERS ENCLOSED IN CONNECTION WITH THIS FILING:**

- ☐ Declaration
- ☐ Power of Attorney: ☐ Separate or ☐ Combined with Declaration
- ☐ Assignment to \_\_\_\_\_ and assignment cover sheet

**CERTIFICATE OF MAILING**  
(37 C.F.R. §1.10)

I hereby certify that this paper (along with any referred to as being attached or enclosed) is being deposited with the United States Postal Service on the date shown below with sufficient postage as 'Express Mail Post Office To Addressee' in an envelope addressed to the Commissioner for Patents, Washington, D.C. 20231.

EL 675944098 US  
Express Mail Label No.

October 3, 2000  
Date of Deposit

Gretchen Dieckmann  
Name of Person Mailing Paper  
*Gretchen Dieckmann*  
Signature of Person Mailing Paper

- ☐ Verified Statement establishing "Small Entity" under 37 CFR §§ 1.9 and 1.27
- ☐ Priority Document No(s):
- ☐ Information Disclosure Statement w/PTO 1449 ☐ Copy of Citations
- ☐ Preliminary Amendment
- ☒ Return Postcard

**III. THE FILING FEE HAS BEEN CALCULATED AS SHOWN BELOW:**

<b>BASIC FILING FEE:</b>				<b>\$690.00</b>
Total Claims	- 20 =	x \$18.00		0.00
Independent Claims	- 3 =	x \$78.00		0.00
Multiple Dependent Claims	\$260 (if applicable)	<input type="checkbox"/>		\$0.00
<b>TOTAL OF ABOVE CALCULATIONS</b>				
Reduction by 1/2 for Filing by Small Entity. Note 37 CFR §§ 1.9, 1.27, 1.28. If applicable, Verified Statement must be attached. <input type="checkbox"/>				
Misc. Filing Fees (Recordation of Assignment -- \$40)				\$0.00
<b>TOTAL FEES SUBMITTED HEREWITH</b>				<b>\$0.00</b>



Customer Number or Bar Code Label

or

New Correspondence Address Below:



PATENT TRADEMARK OFFICE

Name					
Address					
City		State		Zip Code	
Country		Telephone		Fax	

**IV. METHOD OF PAYMENT OF FEES**

- ☐ A check in the amount of \$\_\_\_\_\_.
- ☒ This application is being filed without fee or Declaration under 37 CFR § 1.53.

Respectfully submitted,

BROBECK, PHLEGER & HARRISON LLP

Dated:

October 3, 2000

By: 

Douglas C. Murdock  
Reg. No. 37,549

12390 El Camino Real  
San Diego, CA 92130  
Telephone: (858) 720-2500  
Facsimile: (858) 720-2555

CONTINUATION-IN-PART APPLICATION

UNDER 37 CFR § 1.53(B)

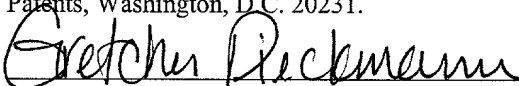
TITLE: G-PROTEIN FUSION RECEPTORS AND  
CHIMERIC GABA<sub>B</sub> RECEPTORS

APPLICANT(S): Thomas Stormann, Lance G. Hammerland, Laura L.  
Storjohann, James G. Busby, James E. Garrett,  
Rachel T. Simin

Correspondence Enclosed:

Continuation-in-Part Transmittal (3 pgs); Cover Sheet  
(1pg); Description (33 pgs); Claims (7 pgs); Abstract (1  
pg); Sequence Listing (102 pgs); Figures (116); and  
Postcard

"EXPRESS MAIL" Mailing Label Number EL675944098US Date of Deposit October 3,  
2000 I hereby certify under 37 CFR §1.10 that this correspondence is being deposited with  
the United States Postal Service as "Express Mail Post Office to Addressee" with sufficient  
postage on the date indicated above and is addressed to the Assistant Commissioner for  
Patents, Washington, D.C. 20231.

  
Gretchen Dieckmann



# G-PROTEIN FUSION RECEPTORS AND CHIMERIC GABA<sub>B</sub> RECEPTORS

## RELATED APPLICATIONS

5           The present application is a continuation in part of PCT/US99/07333 which claims priority to Garrett *et al.* U.S. Serial No. 60/080,671, filed April 3, 1998, which is hereby incorporated by reference herein in its entirety including the drawings.

## FIELD OF THE INVENTION

10           The present invention relates to a G-protein fusion receptors, chimeric GABA<sub>B</sub> ( $\gamma$ -aminobutyric acid) receptors, nucleic acid encoding such receptors, and uses of such receptors and nucleic acid encoding such receptors.

## BACKGROUND

15           The references cited herein are not admitted to be prior art to the claimed invention.

          Chimeric receptors made up of peptide segments from different receptors have different uses such as being used to assess the functions of different sequence regions and to assess the activity of different compounds at a particular receptor. Examples of using  
20           chimeric receptors to assess the activity of different compounds are provided by Dull *et al.*, U.S. Patent No. 4,859,609, Dull *et al.*, U.S. Patent No. 5,030,576, and Fuller *et al.*, U.S. Patent No. 5,981,195.

          Dull *et al.* U.S. Patent No. 4,859,609, and Dull *et al.* U.S. Patent No. 5,030,576, indicate the production and use of chimeric receptors comprising a ligand binding domain  
25           of a predetermined receptor and a heterologous reporter polypeptide. The Dull *et al.* patents provide as examples of chimerics: (1) a chimeric receptor made up of the insulin receptor extracellular chain, and the EGF receptor transmembrane and cytoplasmic domains without any HIR B-chain sequence; and (2) a hybrid receptor made up of the v-  
erB oncogene product intracellular domain fused to the EGF receptor extracellular and  
30           transmembrane domains.

          Fuller *et al.* International Publication No. WO 97/05252 feature chimeric receptors made up of metabotropic glutamate receptor (mGluR) domains and calcium receptor

(CaR) domains. The chimeric receptors allow the coupling of functional aspects of a mGluR with a CaR.

An example of the use of chimeric receptors to assess the functions of different sequence regions receptors are found in studies identifying regions of different guanine nucleotide-binding protein coupled receptors important for guanine nucleotide-binding protein coupling. (See, Kobilka *et al.*, *Science* 240:1310-1316, 1988; Wess *et al.*, *FEBS Lett.* 258:133-136, 1989; Cotecchia *et al.*, *Proc. Natl. Acad. Sci. USA* 87:2896-2900, 1990; Lechleiter *et al.*, *EMBO J.* 9:4381-4390, 1990; Wess *et al.*, *Mol. Pharmacol.* 38:517-523, 1990; and Pin *et al.*, *EMBO J.* 13:342-348, 1994.)

### SUMMARY OF THE INVENTION

The present invention features G-protein fusion receptors and chimeric GABA<sub>B</sub> receptors (GABA<sub>B</sub>Rs), nucleic acid encoding such receptors, and the use of such receptors and nucleic acid. G-protein fusion receptors comprise at least one domain from a CaR, a mGluR, and/or a GABA<sub>B</sub> receptor fused directly or through a linker to a guanine nucleotide-binding protein (G-protein). Chimeric GABA<sub>B</sub>Rs comprise at least one of a GABA<sub>B</sub>R extracellular domain, a GABA<sub>B</sub>R transmembrane domain, or a GABA<sub>B</sub>R intracellular domain and one or more domains from a mGluR subtype 8 (mGluR8) and/or a CaR.

G-proteins are peripheral membrane proteins made up of an  $\alpha$  subunit, a  $\beta$  subunit, and a  $\gamma$  subunit. G-proteins interconvert between a GDP bound and a GTP bound form. Different types of G-proteins can affect different enzymes, such as adenylate cyclase and phospholipase-C.

Thus, a first aspect of the present invention describes a G-protein fusion receptor comprising:

an extracellular domain comprising an amino acid sequence substantially similar to either an extracellular CaR amino acid sequence, an extracellular mGluR amino acid sequence, or an extracellular GABA<sub>B</sub> receptor amino acid sequence;

a transmembrane domain joined to the carboxy terminus of said extracellular domain, said transmembrane domain comprising a transmembrane domain amino acid sequence substantially similar to either a transmembrane CaR amino acid sequence, a transmembrane mGluR amino acid sequence, or a transmembrane GABA<sub>B</sub> receptor amino acid sequence;

an intracellular domain joined to the carboxy terminus of said transmembrane domain comprising all or a portion of an intracellular amino acid sequence substantially similar to either an intracellular CaR amino acid sequence, an intracellular mGluR amino acid sequence, or an intracellular GABA<sub>B</sub> receptor amino acid sequence, provided that  
5 said portion is at least about 10 amino acids;

an optionally present linker joined to the carboxy terminus of said intracellular domain, where said optionally present linker is a polypeptide 3 to 30 amino acids in length, wherein said amino acids of the optionally present linker are selected from the group consisting of alanine, proline, serine, and glycine; and

10 a G-protein joined either to said intracellular domain or to said optionally present linker, provided that said G-protein is joined to said optionally present linker when said optionally present linker is present.

“Substantially similar” refers to at least 40% sequence similarity between respective polypeptide regions making up a domain. In preferred embodiments,  
15 substantially similar refers to at least 50%, at least 75%, at least 90%, at least 95% sequence similarity, or 100% (the same sequence), between polypeptide domains. The degree to which two polypeptide domains are substantially similar is determined by comparing the amino acid sequences located in corresponding domains. Sequence similarity is preferably determined using BLASTN (Altschul *et al.*, *J. Mol. Biol.* 215:403-  
20 410, 1990).

The different receptor components of the G-protein receptor can come from the same receptor protein or from a chimeric receptor made up of different receptor domains. By swapping different domains compounds able to effect different domains of a particular receptor can be identified and the activity of different compounds at different  
25 domains can be measured.

In different embodiments the CaR region(s) present in the G-protein fusion are substantially similar to, comprise, or consist of portion(s) of a mammalian CaR, preferably the human CaR; mGluR region(s) present in the G-protein fusion are substantially similar to, comprise, or consist of portion(s) of a mammalian mGluR,  
30 preferably a human mGluR; and GABA<sub>B</sub>R region(s) present in the G-protein fusion are substantially similar to, comprise, or consist of portion(s) of a mammalian GABA<sub>B</sub>R, preferably a human GABA<sub>B</sub>R.

In preferred embodiments concerning GABA<sub>B</sub>R regions that are present: the GABA<sub>B</sub>R extracellular domain is substantially similar to a GABA<sub>B</sub>R extracellular domain provided in SEQ. ID. NOs. 2-4; the GABA<sub>B</sub>R transmembrane domain is substantially similar to the GABA<sub>B</sub>R transmembrane domain provided in SEQ. ID. NOs. 7-9; and the GABA<sub>B</sub>R intracellular domain is substantially similar to a GABA<sub>B</sub>R intracellular domain provided in SEQ. ID. NOs. 12-14.

In preferred embodiments concerning CaR regions that are present: the CaR extracellular domain is substantially similar to the CaR extracellular provided in SEQ. ID. NO. 1; the CaR transmembrane domain is substantially similar to the CaR transmembrane domain provided in SEQ. ID. NO. 6; and the CaR intracellular domain is substantially similar to the CaR intracellular domain such as that provided in SEQ. ID. NO. 11.

Various different mGluR subtypes present in different organisms, including humans, are described in different patent publications as follows: mGluR<sub>1</sub> - WO 94/29449, EP 569 240 A1, WO 92/10583 and U.S. Patent No. 5,385,831; mGluR<sub>2</sub> - WO 94/29449, WO 96/06167, and EP 711 832 A2; mGluR<sub>3</sub> - WO 94/29449, and WO 95/22609; mGluR<sub>4</sub> - WO 95/08627, WO 95/22609, and WO 96/29404; mGluR<sub>5</sub> - WO 94/29449; mGluR<sub>6</sub> - WO 95/08627; mGluR<sub>7</sub> - U.S. Patent No. 5,831,047, WO 95/08627 and WO 96/29404; and mGluR<sub>8</sub> - U.S. Patent Nos. 6,051,688, 6,077,675, 6,084,084 and EP 816 498 A2. (Each of these references are hereby incorporated by reference herein.)

In preferred embodiments concerning mGluR regions that are present: the mGluR extracellular domain is substantially similar to either human mGluR 1, human mGluR 2, human mGluR 3, human mGluR 4, human mGluR 5, human mGluR 6, human mGluR 7, or human mGluR 8; the mGluR transmembrane domain is substantially similar to either human mGluR 1, human mGluR 2, human mGluR 3, human mGluR 4, human mGluR 5, human mGluR 6, human mGluR 7, or human mGluR 8; and the mGluR intracellular domain is substantially similar to either human mGluR 1, human mGluR 2, human mGluR 3, human mGluR 4, human mGluR 5, human mGluR 6, human mGluR 7, or human mGluR 8. Preferred embodiments also include any mGluR splice variant.

In preferred embodiments concerning the optionally present linker, said optionally present linker is a polypeptide 3 to 30 amino acids in length, wherein said amino acids of the optionally present linker are selected from the group consisting of alanine, proline, serine, and glycine; and more preferably, the optionally present linker is comprised of alanine amino acids.

Another aspect of the present invention describes a recombinant cell comprising an expression vector encoding for a G-protein fusion receptor, and a cell where the G-protein fusion receptor is expressed. Preferably, the G-protein fusion receptor is functional in the cell.

5 Another aspect of the present invention describes a recombinant cell produced by combining (a) a cell where a G-protein fusion receptor is expressed, and (b) a vector comprising nucleic acid encoding a G-protein fusion receptor and elements for introducing heterologous nucleic acid into the cell. Preferably, the G-protein fusion receptor is functional in the cell.

10 Another aspect of the present invention describes a process for the production of a G-protein fusion receptor. The process is performed by growing host cells comprising a G-protein fusion receptor.

Another aspect of the present invention describes a method of measuring the ability of a compound to affect G-protein fusion receptor activity.

15 Another aspect of the present invention describes a chimeric GABA<sub>B</sub>R comprising an extracellular domain, a transmembrane domain and an intracellular domain, wherein at least one domain is from a GABA<sub>B</sub>R and at least one domain is from CaR or mGluR8. The extracellular domain comprises an amino acid sequence substantially similar to a CaR extracellular domain (SEQ. ID. NO. 1), a GABA<sub>B</sub>R1a extracellular domain (SEQ. ID. NO. 2), a GABA<sub>B</sub>R1b extracellular domain (SEQ. ID. NO. 3), a GABA<sub>B</sub>R2  
20 extracellular domain (SEQ. ID. NO. 4), or a mGluR8 extracellular domain (SEQ. ID. NO. 5).

The transmembrane domain comprises an amino acid sequence substantially similar to a CaR transmembrane domain (SEQ. ID. NO. 6), a GABA<sub>B</sub>R1a transmembrane  
25 domain (SEQ. ID. NO. 7), a GABA<sub>B</sub>R1b transmembrane domain (SEQ. ID. NO. 8), a GABA<sub>B</sub>R2 transmembrane domain (SEQ. ID. NO. 9), or a mGluR8 transmembrane domain (SEQ. ID. NO. 10).

The intracellular domain comprises an amino acid sequence substantially similar to a CaR intracellular domain (SEQ. ID. NO. 11), a GABA<sub>B</sub>R1a intracellular domain  
30 (SEQ. ID. NO. 12), a GABA<sub>B</sub>R1b intracellular domain (SEQ. ID. NO. 13), a GABA<sub>B</sub>R2 intracellular domain (SEQ. ID. NO. 14), or a mGluR8 intracellular domain (SEQ. ID. NO. 15).

Preferred chimeric GABA<sub>B</sub>Rs contain at least one mGluR8 intracellular, transmembrane or extracellular domain, or at least one CaR intracellular, transmembrane or extracellular domain. More preferably, the chimeric GABA<sub>B</sub>R contains at least one CaR domain.

5 In preferred embodiments concerning mGluR8 regions that are present: the mGluR8 extracellular domain is substantially similar to the mGluR8 extracellular domain provided in SEQ. ID. NO. 5; the mGluR8 transmembrane domain is substantially similar to the mGluR8 transmembrane domain provided in SEQ. ID. NO. 10; and the mGluR8 intracellular domain is substantially similar to the mGluR8 receptor intracellular provided  
10 in SEQ. ID. NO. 15.

Preferably, the domains are functionally coupled such that a signal from the binding of an extracellular ligand is transduced to the intracellular domain when the chimeric receptor is present in a suitable host cell. A suitable host cell contains the elements for functional signal transduction for receptors coupled to a G-protein.

15 Another aspect of the present invention describes a nucleic acid comprising a nucleotide sequence encoding for a chimeric GABA<sub>B</sub>R.

Another aspect of the present invention describes a recombinant cell comprising an expression vector encoding for a chimeric GABA<sub>B</sub>R, and a cell where the chimeric GABA<sub>B</sub>R is expressed. Preferably, the chimeric GABA<sub>B</sub>R is functional in the cell.

20 Another aspect of the present invention describes a recombinant cell produced by combining (a) a cell where a chimeric GABA<sub>B</sub>R is expressed, and (b) a vector comprising nucleic acid encoding the chimeric GABA<sub>B</sub>R and elements for introducing heterologous nucleic acid into the cell. Preferably, the chimeric GABA<sub>B</sub>R is functional in the cell.

Another aspect of the present invention describes a process for the production of a  
25 chimeric receptor. The process is performed by growing host cells comprising a chimeric GABA<sub>B</sub>R.

Another aspect of the present invention describes a method of measuring the ability of a compound to affect GABA<sub>B</sub>R or mGluR activity. The method is performed by measuring the ability of a compound to affect chimeric GABA<sub>B</sub>R or mGluR activity.

30 Another aspect of the present invention describes a fusion receptor polypeptide comprising a receptor and a G-protein  $\alpha$  subunit, wherein said G-protein  $\alpha$  subunit is fused to the intracellular domain of said receptor, provided that the receptor is not an adrenoreceptor.

Various examples are described herein. These examples are not intended in any way to limit the claimed invention.

Other features and advantages of the invention will be apparent from the following drawings, the description of the invention, the examples, and the claims.

5

### BRIEF DESCRIPTION OF DRAWINGS

Figures 1a-1d illustrate the amino acid sequences of a human CaR extracellular domain (SEQ. ID. NO. 1), a human GABA<sub>B</sub>R1a extracellular domain (SEQ. ID. NO. 2), a human GABA<sub>B</sub>R1b extracellular domain (SEQ. ID. NO. 3), a human GABA<sub>B</sub>R2  
10 extracellular domain (SEQ. ID. NO. 4), and a human mGluR8 extracellular domain (SEQ. ID. NO. 5).

Figures 2a-2b illustrate the amino acid sequences of a human CaR transmembrane domain (SEQ. ID. NO. 6), a human GABA<sub>B</sub>R1a transmembrane domain (SEQ. ID. NO. 7), a human GABA<sub>B</sub>R1b transmembrane domain (SEQ. ID. NO. 8), a human GABA<sub>B</sub>R2  
15 transmembrane domain (SEQ. ID. NO. 9), and a human mGluR8 transmembrane domain (SEQ. ID. NO. 10).

Figures 3a-3b illustrate the amino acid sequences of a human CaR intracellular domain (SEQ. ID. NO. 11), a human GABA<sub>B</sub>R1a intracellular domain (SEQ. ID. NO. 12), a human GABA<sub>B</sub>R1b intracellular domain (SEQ. ID. NO. 13), a human GABA<sub>B</sub>R2  
20 intracellular domain (SEQ. ID. NO. 14), and a human mGluR8 intracellular domain (SEQ. ID. NO. 15).

Figures 4a-4b illustrate the amino acid sequence of G<sub>15</sub> (SEQ. ID. NO. 16) and G<sub>16</sub> (SEQ. ID. NO. 17).

Figures 5a-5r illustrate the cDNA sequences encoding for human CaR (SEQ. ID. NO. 18), human GABA<sub>B</sub>R1a (SEQ. ID. NO. 19), human GABA<sub>B</sub>R1b (SEQ. ID. NO. 20),  
25 and human GABA<sub>B</sub>R2 (SEQ. ID. NO. 21).

Figures 6a-6h illustrate the cDNA sequence for rat GABA<sub>B</sub>R1a (SEQ. ID. NO. 22) and rat GABA<sub>B</sub>R1b (SEQ. ID. NO. 23).

Figures 7a-7c illustrate the amino sequence for rat GABA<sub>B</sub>R1a (SEQ. ID. NO. 24) and rat GABA<sub>B</sub>R1b (SEQ. ID. NO. 25).  
30

Figure 8 illustrates the ability of a chimeric CaR/GABA<sub>B</sub>R2 (CaR extracellular and transmembrane domains, and intracellular GABA<sub>B</sub>R2 domain) to transduce a signal. Signal production was measured by detecting an increase in the calcium-activated

chloride current. The line in the middle of the increase signifies a wash step.

Figures 9a-9p illustrate the cDNA sequence for human mGluR2 (SEQ. ID. NO. 26), chimeric hCAR/hmGluR2 (SEQ. ID. NO. 30), chimeric hmGluR2/hCaR (SEQ. ID. NO. 34), and chimeric hmGluR8/hCaR (SEQ. ID. NO. 38).

Figures 10a-10f illustrate the amino acid sequence for human mGluR2 (SEQ. ID. NO. 27), chimeric hCAR/hmGluR2 (SEQ. ID. NO. 31), chimeric hmGluR2/hCaR (SEQ. ID. NO. 35), chimeric hmGluR8/hCaR (SEQ. ID. NO. 39).

Figures 11a-11v illustrate the cDNA sequence for the phCaR/hmGluR2\*Gqi5 fusion construct (SEQ. ID. NO. 32), pmGluR2//CaR\*G<sub>qi5</sub> fusion construct (SEQ. ID. NO. 36), pmGluR2//CaR\*G<sub>qi5</sub>+3Ala linker fusion construct (SEQ. ID. NO. 46), and the mGluR8//CaR\*G<sub>qi5</sub> fusion construct (SEQ. ID. NO. 40).

Figures 12a-12h illustrate the amino acid sequence for the phCaR/hmGluR2\*Gqi5 fusion construct (SEQ. ID. NO. 33), pmGluR2//CaR\*G<sub>qi5</sub> fusion construct (SEQ. ID. NO. 37), pmGluR2//CaR\*G<sub>qi5</sub>+3Ala linker fusion construct (SEQ. ID. NO. 47), and the mGluR8//CaR\*G<sub>qi5</sub> fusion construct (SEQ. ID. NO. 41).

Figures 13a-13m illustrate the cDNA sequence for the GABA-R2\*Gqo5 fusion construct (SEQ. ID. NO. 42) and the GABA-BR1a\*Gqo5 fusion construct (SEQ. ID. NO. 44).

Figures 14a-14e illustrates the amino acid sequence for the GABA-BR2\*Gqo5 fusion construct (SEQ. ID. NO. 43) and the GABA-BR1a\*Gqo5 fusion construct (SEQ. ID. NO. 45).

Figure 15 illustrates the ability of different G-protein fusions to transduce signal resulting from ligand binding. mGluR2//CaR\*Gqi5 is shown by SEQ. ID. NO. 37, CaR/mGluR2\*Gqi5 is shown by SEQ. ID. NO. 33, mGluR8//CaR\*Gqi5 is shown by SEQ. ID. NO. 41.

Figures 16a-16e illustrates the amino acid sequence for the ph8SPmGluR4 chimeric construct (SEQ. ID. NO.48), the amino acid sequence for the phmGluR4//CaR\*AAA\*G $\alpha$ <sub>qi5</sub> fusion construct (SEQ. ID. NO. 49), and the phmGluR8//CaR\*AAA\*G $\alpha$ <sub>qi5</sub> fusion construct (SEQ. ID. NO. 50).

## DETAILED DESCRIPTION OF THE INVENTION



The CaR, mGluR, and the GABA<sub>B</sub>R are structurally similar in that they are each a single subunit membrane protein possessing an extracellular domain, a transmembrane domain comprising seven putative membrane spanning helices connected by three intracellular and three extracellular loops, and an intracellular carboxy-terminal domain.

5 Signal transduction is activated by the extracellular binding of an agonist. The signal is transduced to the intracellular components of the receptor causing an intracellular effect.

Signal transduction from agonist binding to an extracellular region can be modulated by compounds acting at a downstream transmembrane domain or the intracellular domain. Downstream effects include antagonist actions of compounds and  
10 allosteric actions of compounds.

The transmembrane domain provides different types of target sites for compounds modulating receptor activity in different environments. As noted above, the transmembrane domain contains extracellular, transmembrane, and intracellular components.

15 Compounds modulating GABA<sub>B</sub>R, CaR, or mGluR activity can be obtained, for example, by screening a group or library of compounds to identify those compounds having the desired activity and then synthesizing such compound. Thus, included in the present invention is a method of making a GABA<sub>B</sub>R, CaR, or mGluR active compound by first screening for a compound having desired properties and then chemically synthesizing  
20 that compound.

### Metabotropic Glutamate Receptors (mGluRs)

mGluRs are G protein-coupled receptors capable of activating a variety of intracellular secondary messenger systems following the binding of glutamate (Schoepp *et al.*, *Trends Pharmacol. Sci.* 11:508, 1990; Schoepp and Conn, *Trends Pharmacol. Sci.* 14:13, 1993, hereby incorporated by reference herein).

Activation of different mGluR subtypes *in situ* elicits one or more of the following responses: activation of phospholipase C, increases in phosphoinositide (PI) hydrolysis, intracellular calcium release, activation of phospholipase D, activation or inhibition of  
30 adenylyl cyclase, increases and decreases in the formation of cyclic adenosine monophosphate (cAMP), activation of guanylyl cyclase, increases in the formation of cyclic guanosine monophosphate (cGMP), activation of phospholipase A<sub>2</sub>, increases in arachidonic acid release, and increases or decreases in the activity of voltage- and ligand-

gated ion channels (Schoepp and Conn, *Trends Pharmacol. Sci.* 14:13, 1993; Schoepp, *Neurochem. Int.* 24:439, 1994; Pin and Duvoisin, *Neuropharmacology* 34:1, 1995, hereby incorporated by reference herein).

Eight distinct mGluR subtypes have been isolated. (Nakanishi, *Neuron* 13:1031, 1994; Pin and Duvoisin, *Neuropharmacology* 34:1, 1995; Knopfel et al., *J. Med. Chem.* 38:1417; *Eur. J. Neuroscience* 7:622-629, 1995, each of these references is hereby incorporated by reference herein.) The different mGluRs possess a large amino-terminal extracellular domain (ECD) followed by a seven putative transmembrane domain (7TMD) comprising seven putative membrane spanning helices connected by three intracellular and three extracellular loops, and an intracellular carboxy-terminal domain of variable length (cytoplasmic tail).

Human mGluR8 is described by Stormann *et al.*, U.S. Patent Nos. 6,051,688, 6,077,675, and 6,084,084, and mouse mGluR8 is described by Duvoisin *et al.*, *J. Neurosci.* 15:3075-3083, 1995, (both of these references are hereby incorporated by reference herein). mGluR8 couples to G<sub>i</sub>. Agonists of mGluR8 include L-glutamate and L-2-amino-4-phosphonobutyrate.

mGluR8 activity can be measured using standard techniques. For example, G<sub>i</sub> negatively couples to adenylate cyclase to inhibit intracellular cAMP accumulation in a pertussis toxin-sensitive fashion. Thus, mGluR8 activity can be measured, for example, by measuring inhibition of forskolin-stimulated cAMP production as described by Duvoisin *et al.*, *J. Neurosci.* 15:3075-3083, 1995.

mGluRs have been implicated in a variety of neurological pathologies. Examples of such pathologies include stroke, head trauma, spinal cord injury, epilepsy, ischemia, hypoglycemia, anoxia, and neurodegenerative diseases such as Alzheimer's disease (Schoepp and Conn, *Trends Pharmacol. Sci.* 14:13, 1993; Cunningham *et al.*, *Life Sci.* 54:135, 1994; Pin et al., *Neuropharmacology* 34:1, 1995; Knopfel et al., *J. Med. Chem.* 38:1417, 1995, each of which is hereby incorporated by reference herein).

### Calcium Receptor

The CaR responds to changes of extracellular calcium concentration and also responds to other divalent and trivalent cations. The CaR is a G-protein coupled receptor containing an extracellular Ca<sup>2+</sup> binding domain. Activation of the CaR, descriptions of CaRs isolated from different sources, and examples of CaR active compound are provided

in Nemeth *NIPS* 10:1-5, 1995, Brown *et al.* U.S. Patent No. 5,688,938, Van Wagenen *et al.*, International Application Number PCT/US97/05558 International Publication Number WO 97/37967, Brown E.M. *et al.*, *Nature* 366:575, 1993, Riccardi D., *et al.*, *Proc. Nat'l. Acad. Sci. USA* 92:131-135, 1995, and Garrett J.E., *et al.*, *J. Biol. Chem.* 31:12919-12925, 1995. (Each of these references are hereby incorporated by reference herein.) Brown *et al.* U.S. Patent No. 5,688,938 and Van Wagenen *et al.*, International Application Number PCT/US97/05558 International Publication Number WO 97/37967, describe different types of compounds active at the CaR including compounds which appear to be allosteric modulators and CaR antagonists.

The CaR can be targeted to achieve therapeutic effects. Examples of target diseases are provided in Brown *et al.* U.S. Patent No. 5,688,938, and Van Wagenen *et al.*, International Application Number PCT/US97/05558 International Publication Number WO 97/37967, and include hyperparathyroidism and osteoporosis.

#### $\gamma$ -Aminobutyric acid Receptors (GABA<sub>B</sub>Rs)

GABA<sub>B</sub>Rs are G-protein coupled metabotropic receptors. GABA<sub>B</sub>Rs modulate synaptic transmission by inhibiting presynaptic transmitter release and by increasing K<sup>+</sup> conductance responsible for long-lasting inhibitory postsynaptic potentials. (See, Kaupmann *et al.*, *Nature* 386:239-246, 1997, hereby incorporated by reference herein.)

GABA<sub>B</sub>Rs are found in the mammalian brain, in locations outside of the brain, and in lower species. Outside of the brain, GABA<sub>B</sub>Rs have been identified on axon terminals and ganglion cell bodies of the autonomic nervous system, on fallopian tube and uterine intestinal smooth muscle cells, in the kidney cortex, urinary bladder muscle and on testicular interstitial cells. (See, Bowery, *Annu. Rev. Pharmacol. Toxicol.* 33:109-147, 1993, hereby incorporated by reference herein.)

Different GABA<sub>B</sub>Rs subtypes exist. Kaupmann *et al.*, *Nature* 386:239-246, 1997, indicate that they cloned GABA<sub>B</sub>Rs. Nucleic acid encoding two GABA<sub>B</sub>R proteins were indicated to be cloned from rat brain: GABA<sub>B</sub>R1a and GABA<sub>B</sub>R1b. GABA<sub>B</sub>R1a differs from GABA<sub>B</sub>R1b in that the N-terminal 147 residues are replaced by 18 amino acids.

GABA<sub>B</sub>R1a and GABA<sub>B</sub>R1b appear to be splice variants. The cloned GABA<sub>B</sub>Rs were indicated to negatively couple adenylyl cyclases and show sequence similarity to the metabotropic receptors for L-glutamate (mGluR). Northern blot analysis indicated that

GABA<sub>B</sub>R1a and GABA<sub>B</sub>R1b is present in brain and testis, but not in kidney, skeletal muscle, liver, lung, spleen, or heart.

Kaupmann *et al.*, International Application Number PCT/EP97/01370, International Publication Number WO 97/46675, indicate that they have obtained rat GABA<sub>B</sub>R clones, GABA<sub>B</sub>R1a and GABA<sub>B</sub>R1b; and humans GABA<sub>B</sub>R clones, GABA<sub>B</sub>R1a/b (representing a partial receptor clone) and GABA<sub>B</sub>R1b (representing a full-length receptor clone). Amino acid sequence information, and encoding cDNA sequence information, is provided for the different GABA<sub>B</sub>R clones.

Another GABA<sub>B</sub>R subtype is GABA<sub>B</sub>R2. Northern blot analysis reveals that an approximately 6.3 Kb human GABA<sub>B</sub>R2 transcript is abundantly expressed in the human brain. Expression is not detected in the heart, placenta, lung, liver, skeletal muscle, kidney and pancreas under conditions where GABA<sub>B</sub>R2 transcript was identified in the human brain. Within the human brain GABA<sub>B</sub>R2 is broadly expressed at variable levels.

GABA<sub>B</sub>R functions as a heterodimer of the subunits GABA<sub>B</sub>R1 or GABA<sub>B</sub>R2. (Jones *et al. Nature* 396:674-679, 1998, hereby incorporated by reference herein.)

GABA<sub>B</sub>Rs have been targeted to achieve therapeutic effects. Kerr and Ong, DDT 1:371-380, 1996, describe different compounds indicated to be GABA<sub>B</sub>R agonists and GABA<sub>B</sub>R antagonists. Kerr and Ong also review therapeutic implications of affecting GABA<sub>B</sub>R activity including, spasticity and motor control, analgesia, epilepsy, cognitive effects, psychiatric disorders, alcohol dependence and withdrawal, feeding behavior, cardiovascular and respiratory functions, and peripheral functions.

Bittiger *et al.*, *Tips* 4:391-394, 1993, review therapeutic applications of GABA<sub>B</sub>R antagonists. Potential therapeutic applications noted by Bittiger *et al.* include cognitive processes, epilepsy, and depression.

### G-Protein Fusion Receptors

Examples of some different types of G-protein fusion receptors, and advantages of some receptors, are provided below. Using the present application as guide additional G-protein receptors fusion can be constructed.

G-protein fusion receptors contain an intracellular domain of a receptor fused to a G-protein subunit (G ). G fusions to adrenoreceptors have been reported by Bertin *et al.*, *Receptors and Channels* 5:41-51, 1997; Wise and Milligan, *Journal of Biological Chemistry* 39:24673-24678, 1997; and Bertin *et al.*, *Proc. Natl. Acad. Sci. USA* 91:8827-

8831, 1994 (each of which are hereby incorporated by reference herein). These studies were indicated to produce a functional chimeric by fusing the  $\alpha_2$ -adrenoreceptor to the  $G_{i1}$ , or the  $\alpha_2$ -adrenoreceptor to the  $G_s$ .

The G-protein fusion receptors described by the present invention include a G-protein fused to an intracellular domain, where the intracellular domain when present in a wild type receptor does not interact with that type of G-protein. Thus, the present invention also describes swapping of signals by fusing an intracellular domain to a G normally not coupled to that intracellular domain. The use of such fusion proteins, while applicable to chimeric GABA<sub>B</sub>Rs, is not limited to chimeric GABA<sub>B</sub>Rs. Indeed, such technology can be applied to receptors containing an extracellular domain, transmembrane domain and intracellular domain of a wild type receptor.

Preferred G-proteins fusion receptors contain an intracellular domain fused to a promiscuous G that couples to phospholipase C resulting in the mobilization of intracellular calcium. Increases in intracellular calcium can be conveniently measured through the use of dyes. Such techniques are well known in the art and are described, for example by Brown *et al.* U.S. Patent No. 5,688,938.

In an embodiment G-proteins fusions can also be used to decrease receptor desensitization.

Examples of promiscuous G's coupling to phospholipase C include naturally occurring G-proteins such as  $G_{15}$  and  $G_{16}$ , and chimeric G-protein such as Gqo5 and Gqi5. Gqo5 and Gqi5 are made of a Gq portion where the five amino acids at the C-terminal are from either  $G_o$  or  $G_i$ , respectively (Conklin *et al.*, *Nature* 363:274-277, 1993, hereby incorporated by reference herein). The Gq portion of such chimeric receptors provides for phospholipase C coupling while the terminal  $G_o$  or  $G_i$  portion allows the chimeric G-protein to couple to different receptor proteins that are normally involved in inhibitor effects on adenylate cyclase.

In an embodiment of the present invention the employed G-protein is from a human source or is made up of different G-protein components each from a human source.

G-proteins fusions can be created, for example, by fusing directly or indirectly the intracellular domain of a receptor protein to a polypeptide having an amino acid sequence substantially similar to  $G_{15}$ ,  $G_{16}$ , Gqo5 or Gqi5. In different embodiments, the receptor

is fused directly or indirectly to a G-protein consisting of the amino acid sequence of G<sub>15</sub>, G<sub>16</sub>, Gqo5 or Gqi5.

The intracellular domain portion of a receptor protein fused directly or indirectly to a G-protein should be at least about 1 amino acid in length. In different embodiments the portion is at least about 10 amino acids, is at least about 50 amino acids, at least about 100 amino acids, or the full length of an intracellular domain.

The intracellular domain can be directly linked to a G-protein or can be indirectly linked through an optionally present linker. Optionally present linkers are preferably about 3 to about 30 amino acids in length. Preferred linkers are made up of alanine, glycine, or a combination thereof.

### Chimeric Receptors

Examples of some different types of chimeric receptors, and advantages of some receptors, are provided below. Using the present application as guide additional chimeric receptors can be constructed.

#### Chimeric GABA<sub>B</sub>R Extracellular Domain

Chimeric GABA<sub>B</sub>Rs containing a GABA<sub>B</sub>R extracellular domain are particularly useful for studying the importance of the GABA<sub>B</sub>R extracellular domain and assaying for compounds active at the extracellular domain. Preferably chimeric GABA<sub>B</sub>Rs containing a GABA<sub>B</sub>R extracellular domain also contain a CaR intracellular domain.

A variety of different activities have been generally attributed to GABA<sub>B</sub>R subtypes. (*E.g.*, Kerr and Ong, DDT 1:371-380, 1996.) Kaupmann *et al.*, *Nature* 386:239-246, 1997, report that in preliminary experiments involving GABA<sub>B</sub>R1a they did not detect positive coupling to the adenylyl cyclase or coupling to the phospholipase effector system.

An intracellular CaR domain can be used to couple with G-proteins which activate phospholipase C and mobilize intracellular calcium. Mobilization of intracellular calcium is readily detected, for example, by fluorescent indicators of intracellular Ca<sup>2+</sup>.

An additional advantage of using the intracellular CaR domain is that CaR G-protein activation is not rapidly desensitized. Thus, the intracellular CaR domain can be used to produce a stronger intracellular signal than a signal produced from a receptor which is desensitized rapidly.

More preferably, the chimeric GABA<sub>B</sub>R contains an intracellular CaR domain, and also contains either a CaR or a GABA<sub>B</sub>R transmembrane domain. Advantages of using a CaR transmembrane domain include separating the effects occurring at a GABA<sub>B</sub>R extracellular domain from effects occurring at a transmembrane domain; and providing additional intracellular elements, present on transmembrane intracellular loops, useful for coupling to G-protein.

A GABA<sub>B</sub>R transmembrane domain is useful for examining whether the transmembrane GABA<sub>B</sub>R can be targeted to affect GABA<sub>B</sub>R activity, and obtaining compounds active at the GABA<sub>B</sub>R transmembrane domain. For example, a transmembrane GABA<sub>B</sub>R can be used to screen for transmembrane allosteric modulators and antagonists.

#### Chimeric GABA<sub>B</sub>R Transmembrane Domain

Chimeric GABA<sub>B</sub>Rs containing a GABA<sub>B</sub>R transmembrane are particularly useful for studying the importance of the GABA<sub>B</sub>R transmembrane domain and assaying for compounds active at the transmembrane domain. Preferably Chimeric GABA<sub>B</sub>Rs containing a GABA<sub>B</sub>R transmembrane domain contain an extracellular domain which is either mGluR8 or CaR, and an intracellular CaR domain.

More preferably, the chimeric GABA<sub>B</sub>R contains an extracellular domain from either mGluR8 or CaR, a GABA<sub>B</sub>R transmembrane, and an intracellular CaR domain. A chimeric GABA<sub>B</sub>R containing extracellular mGluR8 or CaR domains can readily be stimulated using mGluR8 or CaR ligands.

#### Chimeric GABA<sub>B</sub>R Intracellular Domain

Chimeric GABA<sub>B</sub>Rs containing a GABA<sub>B</sub>R intracellular domain are particularly useful for studying the importance of the GABA<sub>B</sub>R intracellular domain and assaying for compounds active at the intracellular domain. Preferably, the chimeric receptors contain an extracellular domain from either mGluR8 or CaR. The extracellular mGluR8 or CaR domains can readily be activated using mGluR8 or CaR ligands.

#### Receptor Domains

Domains of a G-protein fusion receptor, a chimeric receptor, and G , substantially similar to a particular sequence can be readily produced using the disclosure provided

herein in conjunction with information well known in the art. Substantially similar sequences can be obtained taking into account sequence information for a particular type of receptor obtained from different sources, different types of amino acids which are to some extent interchangeable, and the ease of experimentation with which functional  
5 receptor activity can be assayed.

Substantially similar sequences includes amino acid alterations such as deletions, substitutions, additions, and amino acid modifications. A "deletion" refers to the absence of one or more amino acid residue(s) in the related polypeptide. An "addition" refers to the presence of one or more amino acid residue(s) in the related polypeptide. Additions  
10 and deletions to a polypeptide may be at the amino terminus, the carboxy terminus, and/or internal. Amino acid "modification" refers to the alteration of a naturally occurring amino acid to produce a non-naturally occurring amino acid. A "substitution" refers to the replacement of one or more amino acid residue(s) by another amino acid residue(s) in the polypeptide. Derivatives can contain different combinations of alterations including more  
15 than one alteration and different types of alterations.

The sequences of polypeptides can be compared from different sources to help identify variable amino acids not essential for receptor activity. For example, Figure 7 provides the rat GABA<sub>B</sub>R1a and GABA<sub>B</sub>R1b amino acid sequences. The rat GABA<sub>B</sub>R1a and GABA<sub>B</sub>R1b amino acid sequences can be compared with the human GABA<sub>B</sub>R1a and  
20 GABA<sub>B</sub>R1b sequences to identify conserved and variable amino acids. Derivatives can then be produced where a variable amino acid is changed, and receptor activity can be readily tested.

Similarly, the amino acid sequences for CaR, mGluR8, and G-proteins from different sources are either known in the art or can readily be obtained. Examples of such  
25 references are provided above.

While the effect of an amino acid change varies depending upon factors such as phosphorylation, glycosylation, intra-chain linkages, tertiary structure, and the role of the amino acid in the active site or a possible allosteric site, it is generally preferred that a substituted amino acid is from the same group as the amino acid being replaced. To some  
30 extent the following groups contain amino acids which are interchangeable: the basic amino acids lysine, arginine, and histidine; the acidic amino acids aspartic and glutamic acids; the neutral polar amino acids serine, threonine, cysteine, glutamine, asparagine and, to a lesser extent, methionine; the nonpolar aliphatic amino acids glycine, alanine, valine,



isoleucine, and leucine (however, because of size, glycine and alanine are more closely related and valine, isoleucine and leucine are more closely related); and the aromatic amino acids phenylalanine, tryptophan, and tyrosine. In addition, although classified in different categories, alanine, glycine, and serine seem to be interchangeable to some extent, and cysteine additionally fits into this group, or may be classified with the polar neutral amino acids.

While proline is a nonpolar neutral amino acid, its replacement represents difficulties because of its effects on conformation. Thus, substitutions by or for proline are not preferred, except when the same or similar conformational results can be obtained.

The conformation conferring properties of proline residues may be obtained if one or more of these is substituted by hydroxyproline (Hyp).

Examples of modified amino acids include the following: altered neutral nonpolar amino acids such as  $\alpha$ -amino acids of the formula  $H_2N(CH_2)_nCOOH$  where  $n$  is 2-6, sarcosine (Sar), tbutylalanine (t-BuAla), t-butylglycine (t-BuGly), N-methyl isoleucine (N-MeIle), and norleucine (Nleu); altered neutral aromatic amino acids such as phenylglycine; altered polar, but neutral amino acids such as citrulline (Cit) and methionine sulfoxide (MSO); altered neutral and nonpolar amino acids such as cyclohexyl alanine (Cha); altered acidic amino acids such as cysteic acid (Cya); and altered basic amino acids such as ornithine (Orn).

Preferred derivatives have one or more amino acid alteration(s) which do not significantly affect the receptor activity of the related receptor protein. In regions of receptor domains not necessary for receptor activity, amino acids may be deleted, added or substituted with less risk of affecting activity. In regions required for receptor activity, amino acid alterations are less preferred as there is a greater risk of affecting receptor activity.

Derivatives can be produced using standard chemical techniques and recombinant nucleic acid techniques. Modifications to a specific polypeptide may be deliberate, as through site-directed mutagenesis and amino acid substitution during solid-phase synthesis, or may be accidental such as through mutations in hosts which produce the polypeptide. Polypeptides including derivatives can be obtained using standard techniques such as those described by Sambrook *et al.*, *Molecular Cloning*, Cold Spring Harbor Laboratory Press (1989). For example, Chapter 15 of Sambrook describes procedures for site-directed mutagenesis of cloned DNA.

### Receptor Nucleic Acid

G-protein fusion and chimeric receptor nucleic acid can be produced based on the information provided herein along with standard recombinant nucleic acid techniques.

5 Examples of references describing recombinant nucleic acid techniques include *Molecular Cloning*, Sambrook *et al.*, Cold Spring Harbor Laboratory Press (1989); and *Current Protocols in Molecular Biology*, Frederick *et al.*, John Wiley & Sons, Inc. (1995).

10 Due to the degeneracy of the genetic code different nucleic acid sequences can encode for a particular polypeptide. Thus, a large number of nucleic acids encoding for a receptor having the same amino acid sequence can be produced.

An embodiment of the present invention uses human nucleic acid encoding for the domains from CaR, GABA<sub>B</sub>R1A, GABA<sub>B</sub>R1B, GABA<sub>B</sub>R2 and/or mGluR8. The amino acid sequences of different domains is provided in Figures 1-3.

### Recombinant Cells

15 Nucleic acid expressing a functional G-Protein fusion or a chimeric receptor can be used to create transfected cells lines expressing such receptors. Such cell lines have a variety of uses such as being used for high-throughput screening for compounds modulating receptor activity; being used to assay binding to the receptor; and as factories to produce large amounts of a receptor.

20 A variety of cell lines can couple exogenously expressed receptors to endogenous functional responses. Cell lines such as NIH-3T3, HeLa, NG115, CHO, HEK 293 and COS7 which are expected to lack CaR, mGluR8, and GABA<sub>B</sub>R can be tested to confirm that they lack these receptors.

25 Production of stable transfectants can be accomplished by transfection of an appropriate cell line with, for example, an expression vector such as pMSG vector, in which the coding sequence for the G-protein fusion or chimeric GABA<sub>B</sub>R cDNA has been cloned. Expression vectors containing a promoter region, such as the mouse mammary tumor virus promoter (MMTV), drive high-level transcription of cDNAs in a variety of mammalian cells. In addition, these vectors contain genes for selecting cells stably expressing cDNA of interest. The selectable marker in the pMSG vectors encode an

30

enzyme, xanthine-guanine phosphoribosyl transferase (XGPRT), conferring resistance to a metabolic inhibitor that is added to the culture to kill nontransfected cells.

The most effective method for transfection of eukaryotic cell lines with plasmid DNA varies with the given cell type. The expression construct will be introduced into cultured cells by the appropriate technique, such as  $\text{Ca}^{2+}$  phosphate precipitation, DEAE-dextran transfection, lipofection or electroporation. Expression of the receptor cDNA in cell lines can be assessed by solution hybridization and Northern blot analysis.

### Binding Assays

The present invention also includes using G-protein fusion receptors or chimeric  $\text{GABA}_\text{B}\text{R}$  in a binding assay. G-protein fusion receptors or chimeric  $\text{GABA}_\text{B}\text{Rs}$  having a particular  $\text{GABA}_\text{B}\text{R}$  domain can be used, for example to facilitate obtaining compounds able to bind to that particular receptor domain; and to determine whether a compound which binds to a particular domain. For example, in a complete chimeric  $\text{GABA}_\text{B}\text{R}$  containing extracellular, transmembrane, and intracellular domains, the presence of one or more domains from  $\text{CaR}$  or  $\text{mGluR}$  are useful to present  $\text{GABA}_\text{B}\text{R}$  domain(s) to a binding agent in a form more like the  $\text{GABA}_\text{B}\text{R}$  domain(s) in the wild type receptor compared to an incomplete  $\text{GABA}_\text{B}\text{R}$  receptor fragment lacking one or more domains.

Binding assays can be carried out using techniques well known in the art. Binding assays preferably employ radiolabeled binding agents.

An example of a binding procedure is carried out by first attaching chimeric  $\text{GABA}_\text{B}\text{R}$  to a solid-phase support to create an affinity matrix. The affinity matrix is then contacted with potential  $\text{GABA}_\text{B}\text{R}$  binding agents. A large library of compounds may be used to determine those compounds binding to the affinity matrix. Bound compounds can be eluted from the column.

### Therapeutic Modulation

As pointed out above, different types of diseases and disorders can be treated using compounds modulating  $\text{CaR}$ ,  $\text{mGluR}$ , or  $\text{GABA}_\text{B}\text{R}$  activity. Additionally, such compounds can be used prophylactically. Compounds modulating  $\text{GABA}_\text{B}\text{R2}$  activity can be administered to patients who would benefit from such treatment. Patients are mammals, preferably humans.

Modulators of CaR, mGluR, or GABA<sub>B</sub>R activity can be administered to a patient using standard techniques. Techniques and formulations generally may be found in Remington's Pharmaceutical Sciences, 18<sup>th</sup> ed., Mack Publishing Co., Easton, PA, 1990 (hereby incorporated by reference herein).

5           Suitable dosage forms, in part, depend upon the use or the route of entry, for example, oral, transdermal, transmucosal, or by injection (parenteral). Such dosage forms should allow the therapeutic agent to reach a target cell whether the target cell is present in a multicellular host or in culture. For example, pharmacological compounds or compositions injected into the blood stream should be soluble. Other factors are well  
10       known in the art, and include considerations such as toxicity and dosage forms which retard the therapeutic agent from exerting its effect.

Therapeutic compounds can be formulated as pharmaceutically acceptable salts and complexes thereof. Pharmaceutically acceptable salts are non-toxic salts in the amounts and concentrations at which they are administered. The preparation of such salts  
15       can facilitate the pharmacological use by altering the physical characteristics of the compound without preventing it from exerting its physiological effect. Useful alterations in physical properties include lowering the melting point to facilitate transmucosal administration and increasing the solubility to facilitate administering higher concentrations of the drug.

20       The pharmaceutically acceptable salt of a compound may be present as a complex. Examples of complexes include an 8-chlorotheophylline complex (analogous to, *e.g.*, dimenhydrinate:diphenhydramine 8-chlorotheophylline (1:1) complex; Dramamine) and various cyclodextrin inclusion complexes.

Pharmaceutically acceptable salts include acid addition salts such as those  
25       containing sulfate, hydrochloride, fumarate, maleate, phosphate, sulfamate, acetate, citrate, lactate, tartrate, methanesulfonate, ethanesulfonate, benzenesulfonate, *p*-toluenesulfonate, cyclohexylsulfamate and quinate.

Pharmaceutically acceptable salts can be obtained from acids such as hydrochloric acid, maleic acid, sulfuric acid, phosphoric acid, sulfamic acid, acetic acid, citric acid,  
30       lactic acid, tartaric acid, malonic acid, methanesulfonic acid, ethanesulfonic acid, benzenesulfonic acid, *p*-toluenesulfonic acid, cyclohexylsulfamic acid, fumaric acid, and quinic acid.

Pharmaceutically acceptable salts also include basic addition salts such as those containing benzathine, chlorprocaine, choline, diethanolamine, ethylenediamine, meglumine, procaine, aluminum, calcium, lithium, magnesium, potassium, sodium, ammonium, alkylamine, and zinc, when acidic functional groups, such as carboxylic acid or phenol are present. For example, see Remington's Pharmaceutical Sciences, 18<sup>th</sup> ed., Mack Publishing Co., Easton, PA, p. 1445, 1990. Such salts can be prepared using the appropriate corresponding bases.

Carriers or excipients can also be used to facilitate administration of therapeutic agents. Examples of carriers include calcium carbonate, calcium phosphate, various sugars such as lactose, glucose, or sucrose, or types of starch, cellulose derivatives, gelatin, vegetable oils, polyethylene glycols and physiologically compatible solvents. Examples of physiologically compatible solvents include sterile solutions of water for injection (WFI), saline solution and dextrose.

GABA<sub>B</sub>R modulating compounds can be administered by different routes including intravenous, intraperitoneal, subcutaneous, intramuscular, oral, topical (transdermal), or transmucosal administration. For systemic administration, oral administration is preferred. For oral administration, for example, the compounds can be formulated into conventional oral dosage forms such as capsules, tablets, and liquid preparations such as syrups, elixirs, and concentrated drops.

Alternatively, injection (parenteral administration) may be used, *e.g.*, intramuscular, intravenous, intraperitoneal, and subcutaneous. For injection, compounds are formulated in liquid solutions, preferably, in physiologically compatible buffers or solutions, such as saline solution, Hank's solution, or Ringer's solution. In addition, the compounds may be formulated in solid form and redissolved or suspended immediately prior to use. Lyophilized forms can also be produced.

Systemic administration can be by transmucosal or transdermal means. For transmucosal or transdermal administration, penetrants appropriate to the barrier to be permeated are used in the formulation. Such penetrants are well known in the art, and include, for example, for transmucosal administration, bile salts and fusidic acid derivatives. In addition, detergents may be used to facilitate permeation. Transmucosal administration, for example, may be through nasal sprays, rectal suppositories, or vaginal suppositories.

For topical administration, compounds can be formulated into ointments, salves, gels, or creams, as is well known in the art.

The amounts of various GABA<sub>B</sub>R modulating compounds to be administered can be determined by standard procedures taking into account factors such as the compound  
5 IC<sub>50</sub>, EC<sub>50</sub>, the biological half-life of the compound, the age, size and weight of the patient, and the disease or disorder associated with the patient. The importance of these and other factors to be considered are well known to those of ordinary skill in the art. Generally, the amount is expected to preferably be between about 0.01 and 50 mg/kg of the animal to be treated.

10

### EXAMPLES

Examples are provided below illustrating different aspects and embodiments of the present invention. The examples include techniques that can be used to produce and use G-protein fusion receptors and chimeric receptors. These examples are not intended to  
15 limit the claimed invention.

#### Example 1: Construction of G-Protein Fusions

This example illustrates different G-protein fusion receptor constructs and techniques used to produce different G-protein fusion receptor constructs. Numbering of  
20 nucleotide position for all the following constructs is such that nucleotide number 1 corresponds to the A of the ATG start codon of the nucleotide sequence encoding the designated protein.

### I. FULL-LENGTH CONSTRUCTS

#### A. phCaR

The cDNA encoding the human CaR (Garrett et al., (1995) J. Biol. Chem.270:12919) is harbored in the Bluescript SK(-) plasmid (Stratagene). This construct is referred to as phCaR.

#### B. phmGluR2

A full length human mGluR2 cDNA was amplified from human cerebellum MarathonReady cDNA (Clontech) using PCR primers based on the human mGluR2 cDNA sequence (Genbank Accession # 4504136). The obtained PCR fragment was

subcloned into the pT7Blue TA vector (Novagen). A Hind III-Not I fragment containing the human mGluR2 cDNA was then subcloned into the Bluescript SKII(-) plasmid (Stratagene). This construct is referred to as phmGluR2.

#### 5 C. phGα<sub>q</sub>

A full length human Gα<sub>q</sub> cDNA was amplified from human cerebral cortex Quick-Clone cDNA (Clontech) using PCR primers based on the human Gα<sub>q</sub> cDNA sequence (Genbank Accession # 4504044). The obtained PCR fragment was subcloned into the Bluescript SKII(-) plasmid (Stratagene). This construct is referred to as phGα<sub>q</sub>.

#### 10 D. phmGluR8

The cDNA encoding the full length human mGluR8 cDNA (Stormann *et al.*, U.S. Patent Nos. 6,051,688, 6,077,675, and 6,084,084) is harbored in the Bluescript SKII(-) plasmid (Stratagene). This construct is referred to as phmGluR8.

#### 15 E. ph8SPmGluR4

A full length human mGluR4 cDNA was amplified from human cerebellum MarathonReady cDNA (Clontech) using PCR primers based on the human mGluR4 cDNA sequence (Genbank Accession #X80818). The obtained PCR fragment was cloned into the pT7Blue TA vector (Novagen). A 2977 bp BamHI fragment containing the human mGluR4 cDNA was then subcloned into the vector pcDNA3.1/Hygro<sup>+</sup> (Invitrogen). This construct is referred to as phmGluR4.

Next, the predicted signal peptide of mGluR4 was replaced with the predicted signal peptide and 87 bp of 5' UTR from phmGluR8 using a recombinant PCR strategy similar to those described above. The first reaction used a phmGluR8 construct with two primers, 3.1-535F (sense 21-mer, complementary to vector sequence upstream of the hmGluR8 insert; sequence 5'-ggcattatgccagtagcatga-3'), and the hybrid primer 8/4RP (antisense 42-mer, containing 21 nucleotides complementary to human mGluR8 and 21 nucleotides complementary human mGluR4; sequence 5'-caagcctctcttcccaggcattttctccacaggtgtattgc-3'). These primers were used to amplify a 469 bp PCR fragment of human mGluR8.

In a separate PCR reaction using phmGluR4 as template, a 472 bp fragment of

human mGluR4 was amplified using a hybrid primer 4/8RP (sense 42-mer, exactly complementary to primer 8/4RP) and oligo mG4-472R, (antisense 18-mer, complementary to the human mGluR4 cDNA; sequence 5'-ctgaagcaccgatgacac-3'). The two PCR products generated from the above two reactions were annealed together in equimolar ratios in the presence of the external primers mG4-472R and 3.1-535F, and Turbo Pfu DNA polymerase (Stratagene).

The resulting chimeric PCR product was digested with NarI and NheI (New England Biolabs) and subcloned into phmGluR4 digested with the same two restriction enzymes. The sequence of the resultant chimeric construct, ph8SPmGluR4, was verified by ABI automated DNA sequence analysis. The replacement of the predicted signal peptide of mGluR4 with that of mGluR8 greatly increased the activity of this receptor in *in vitro* assays

## II. G $\alpha$ q15

The cDNA encoding the human G $\alpha$ q15 cDNA (Conklin et al (1993) Nature 363:274-77) is harbored in the Bluescript SKII(-) plasmid (Stratagene). This construct is referred to as G $\alpha$ q15. The nucleic acid and amino acid sequences for G $\alpha$ q15 are provided by SEQ. ID. NOs. 28 and 29 respectively.

## III. phCaR/hmGluR2

This chimera contains the extracellular domain of the human CaR and transmembrane domain and intracellular cytoplasmic tail of human mGluR2. The chimeric junction between the CaR and hmGluR2 was created using a recombinant PCR strategy similar to those described above.

The first reaction used two primers, CA1156 (sense 19-mer, corresponding to nucleotides 1156-1174 of human CaR), and the hybrid primer CA/2 (antisense 42-mer, containing 21 nucleotides complementary to nucleotides 1774-1794 of human CaR and 21 nucleotides complementary to nucleotides 1660-1680 of the human mGluR2). These primers were used to amplify a 659 bp PCR fragment of human CaR.

In a separate PCR reaction using phmGluR2 as template, a 692 bp fragment of the human mGluR2 was amplified using a hybrid primer 2/CA (sense 42-mer, exactly complementary to primer CA/2) and oligo 2-2330m, (antisense 23-mer, complementary to



nucleotides 2309-2331 of the human mGluR2 cDNA). The two PCR products generated from the above two reactions were annealed together in equimolar ratios in the presence of the external primers CA1156 and 2-2330m, and the Pfu DNA polymerase (Stratagene).

5 The resulting chimeric PCR product was digested with SexA1 (Boehringer Mannheim) and BamHI (New England Biolabs) and subcloned into phCaR digested with the same two restriction enzymes. In the final cloning step, the 3' end of human mGluR2 was subcloned into this construct using the restriction enzymes BsrGI and BamHI (both New England Biolabs). The sequence of the resultant chimeric construct,  
10 phCaR/hmGluR2, was verified by ABI automated DNA sequence analysis.

#### IV. phCaR/hmGluR2\*Gqi5

This construct contains the phCaR/hmGluR2 chimeric receptor fused to human  $G\alpha_{qi5}$ . A HindIII-BamHI fragment containing the phCaR/hmGluR2 construct was  
15 subcloned into pcDNA3.1/Hygro(+) (Invitrogen) to aid in constructing this fusion protein.

The chimeric junction between the C-terminus of phCaR/hmGluR2 and the N-terminus of  $G\alpha_{qi5}$  was created using a recombinant PCR strategy similar to those described above.

The first reaction used two primers, 2-1713 (sense 21-mer, corresponding to  
20 nucleotides 1710-1730 of human mGluR2) and the hybrid primer 2/Q (antisense 42-mer, containing 21 nucleotides complementary to nucleotides 2596-2616 of human mGluR2, and 21 nucleotides complementary to nucleotides 1-21 of p $G\alpha_{qi5}$ ). These primers were used to amplify a 927 bp PCR fragment of phCaR/hmGluR2. In a separate PCR reaction all of  $G\alpha_{qi5}$  was amplified using a hybrid primer Q/2 (sense 42-mer, exactly  
25 complementary to primer 2/Q) and the and the T3 primer commercially available from Stratagene.

These two PCR products generated from the above two reactions were annealed together in equimolar ratios in the presence of the external primers 2-1713 and T3, and the Pfu DNA polymerase (Stratagene). The resulting chimeric PCR product was digested  
30 with Bsu361 and BamHI (New England Biolabs) and subcloned into phCaR/hmGluR2 digested with the same two restriction enzymes. The sequence of the resultant chimeric fusion construct, phCaR/hmGluR2\* $G\alpha_{qi5}$ , was verified by DNA sequence analysis.

### V. phmGluR2//CaR Construct

This chimera contains the extracellular and transmembrane domains of human mGluR2 linked to the intracellular cytoplasmic tail domain of the human CaR. The chimeric junction was created using three separate PCR reactions.

The first reaction used two primers, 2-1713 (sense 21-mer, corresponding to nucleotides 1710-1730 of human mGluR2, Genbank Accession # 4504136) and the hybrid primer 2/CT (antisense 42-mer, containing 21 nucleotides complementary to nucleotides 2452 – 2472 of human mGluR2 and 21 nucleotides complementary to nucleotides 2602-2622 of the human CaR). These primers were used to amplify a 783 bp PCR fragment of human mGluR2. In a separate PCR reaction using phCaR in the BlueScript SK<sup>-</sup> plasmid as template, a 750 bp fragment of the human CaR was amplified using a hybrid primer CT/2 (sense 42-mer, exactly complementary to primer 2/CT) and the T3 primer commercially available from Stratagene.

The two PCR products generated from the above two reactions were annealed together in equimolar ratios in the presence of the external primers 2-1713 and T3, and the Pfu DNA polymerase (Stratagene). The resulting chimeric PCR product was digested with BsrG I and Not I (New England Biolabs) and subcloned into pmGluR2 digested with the same two restriction enzymes. The sequence of the resultant chimeric construct, pmGluR2//CaR, was verified by ABI automated DNA sequence analysis.

### VI. pmGluR2//CaR\*Gα<sub>q</sub>i5 Construct

This construct contains the hmGluR2//CaR chimeric receptor fused to human Gα<sub>q</sub>i5. The chimeric junction between the C-terminus of hmGluR2//CaR and the N-terminus of Gα<sub>q</sub>i5 was created using a recombinant PCR strategy similar to that described above for the construction of phmGluR2//CaR.

The first reaction used two primers, CRP10A (sense 18-mer, corresponding to nucleotides 2812-2829 of phCaR) and the hybrid primer CaRQ (antisense 42-mer, containing 21 nucleotides complementary to nucleotides 3214– 3234 phCaR, and 21 nucleotides complementary to nucleotides 1-21 of pGα<sub>q</sub>i5). These primers were used to amplify a 443 bp PCR fragment of hmGluR2//CaR. In a separate PCR reaction, all of Gα<sub>q</sub>i5 was amplified using a hybrid primer QCaR (sense 42-mer, exactly complementary

to primer CaRQ) and the T3 primer commercially available from Stratagene.

The two PCR products generated from the above two reactions were annealed together in equimolar ratios in the presence of the external primers CRP10A and T3, and the Pfu DNA polymerase (Stratagene). The resulting chimeric PCR product was digested with BstE II and Not I (New England Biolabs) and subcloned into pmGluR2//CaR  
 5 digested with the same two restriction enzymes. The sequence of the resultant chimeric fusion construct, pmGluR2//CaR\*G $\alpha$ <sub>q</sub>i5, was verified by ABI automated DNA sequence analysis.

## VII. Fusion Receptor Protein Linker Addition Constructs

### A. phmGluR2//CaR\*AAA\*G $\alpha$ <sub>q</sub>i5

A linker encoding three alanine residues was incorporated into the phmGluR2//CaR\*G $\alpha$ <sub>q</sub>i5 construct by mutagenesis (Stratagene QuickChange Mutagenesis  
 15 Kit). A sense 40-mer, 2CQ+LP, contained 16 nucleotides corresponding to 3219-3234 of human CaR, followed by the 9 nucleotide sequence (GCGGCCGCC) encoding three alanine residues and a NotI restriction enzyme site, and then 15 nucleotides corresponding to nucleotides 1-15 of G $\alpha$ <sub>q</sub>i5. 2CQ+LP was annealed to an antisense 40-mer, 2CQ+LM, the exact complement of 2CQ+LP. These oligos were used in the mutagenesis reaction  
 20 according to the manufacturer's protocol. Restriction enzyme analysis and DNA sequence analysis confirmed the insertion of the 9 nucleotide linker (GCGGCCGCC) between the C-terminus of phmGluR2//CaR and the N-terminus of G $\alpha$ <sub>q</sub>i5. This construct was designated phmGluR2//CaR\*AAA\*G $\alpha$ <sub>q</sub>i5.

### B. Human GABA<sub>B</sub>R2\*AAA\*G $\alpha$ <sub>q</sub>o5 and human GABA<sub>B</sub>R1a\*AAA\*G $\alpha$ <sub>q</sub>o5

These constructs contain the human GABA<sub>B</sub>R2 (hGABA<sub>B</sub>R2: Genbank Accession # AJ 012188) and human GABA<sub>B</sub>R1a (hGABA<sub>B</sub>R1a: Genbank Accession # AJ 012185) fused at their C-terminus to the N-terminus of human G $\alpha$ <sub>q</sub>o5 (hG $\alpha$ <sub>q</sub>o5: *Nature* 363:274-276, 1993). Human GABA<sub>B</sub>R2, hGABA<sub>B</sub>R1a, and hG $\alpha$ <sub>q</sub>o5 were cloned into the plasmid  
 30 pcDNA3.1/Hygro+ (Invitrogen) and are designated phGABA<sub>B</sub>R2, phGABA<sub>B</sub>R1a, and phG $\alpha$ <sub>q</sub>o5. The first reaction used two primers, XcmI-R2 (sense 20-mer, corresponding to nucleotides 2650-2669 of phGABA<sub>B</sub>R2) and the hybrid primer R2/Go5(-) (antisense 45-

mer, containing 18 nucleotides complementary to nucleotides 2806-2823 of phGABA<sub>B</sub>R2 and 18 nucleotides complementary to nucleotides 1-18 of hGα<sub>q</sub>o5). These two complementary areas flank a 9 nucleotide sequence coding for 3 alanine sequences with a unique NotI restriction site. These primers were used to amplify a 200 base-pair PCR  
 5 fragment.

In a separate PCR reaction, part of hGα<sub>q</sub>o5 was amplified using a hybrid primer R2/Gα<sub>q</sub>o5(+) (sense 45-mer), exactly complementary to R2/Go5(-) and XbaI-Go5 primer (22-mer containing 22 nucleotides complementary to nucleotides 873-895 of hGα<sub>q</sub>o5). These primers were used to amplify a 914 base-pair PCR product. The two PCR products  
 10 generated from the above two reactions were annealed together in equimolar ratios in the presence of the external primers; XcmI-R2 and XbaI-Go5, and Pfu polymerase (Stratagene).

The resulting chimeric PCR product was digested with the restriction endonucleases XcmI and XbaI (New England Biolabs) and subcloned into phGABA<sub>B</sub>R2 digested with the same two restriction enzymes. The resulting clone was then digested  
 15 with HindIII and XbaI and subcloned into phGα<sub>q</sub>o5 cut with HindIII and XbaI resulting in the chimeric hGABA<sub>B</sub>R\*AAA\*Gα<sub>q</sub>o5. The chimeric junction between the C-terminus hGABA<sub>B</sub>R1a, the Ala linker, and the N-terminus of hGα<sub>q</sub>o5 was created using a recombinant PCR strategy similar to those described above.

To construct hGABA<sub>B</sub>R1a\*AAA\*Gqo5, the first reaction used a commercially available T7 primer (Novagen) and the Ntl hGBR1 primer  
 20 (CAGAGTCATGGCGGCCGCCTTATAAAGCAAATGCACTCG) corresponding to nucleotide numbers 1-9 of hGα<sub>q</sub>o5 and nucleotide numbers 2863-2883 of hGABA<sub>B</sub>R1a.

### 25 C. phmGluR8//CaR\*AAA\*Gα<sub>q</sub>i5

A linker encoding three alanine residues was incorporated into the phmGluR8//CaR\*Gα<sub>q</sub>i5 construct by mutagenesis (Stratagene QuickChange Mutagenesis Kit), exactly as described in Section A, above to create phmGluR2//CaR\*AAA\*Gα<sub>q</sub>i5. The same primers, 2CQ+LP and 2CQ+LM, were used for this mutagenesis. Restriction  
 30 enzyme analysis and DNA sequence analysis confirmed the insertion of the 9-nucleotide linker (GCGGCCGCC) between the C-terminus of phmGluR8//CaR and the N-terminus of Gα<sub>q</sub>i5. This construct was designated phmGluR8//CaR\*AAA\*Gα<sub>q</sub>i5.

#### D. ph8SPmGluR4//CaR\*AAA\*G $\alpha$ <sub>q</sub>i5

This chimera contains the extracellular and transmembrane domains of the human 8SPmGluR4 construct and intracellular cytoplasmic tail of human CaR fused to G $\alpha$ <sub>q</sub>i5 through the three alanine residue linker.

The chimeric junction between the human 8SPmGluR4 and hCaR was created using a recombinant PCR strategy similar to those previously described. The first reaction used two primers, mG4-2028R (sense 19-mer, corresponding to nucleotides of human 8SPmGluR4; sequence 5'-catctaccgcatcttcgag-3'), and the hybrid primer 4CT (antisense 42-mer, containing 21 nucleotides complementary to human 8SPmGluR4 and 21 nucleotides complementary human CaR; sequence 5'-acgcacctcctcgatgggtgttctgctccgggtggaaggagat-3'). These primers were used to amplify a 549 bp PCR fragment from human 8SPmGluR4.

In a separate PCR reaction, using phmGluR2//CaR\*AAA\*G $\alpha$ <sub>q</sub>i5 as a template, a 743 bp fragment of the human CaR\*AAA\*G $\alpha$ <sub>q</sub>i5 was amplified using the hybrid primer CT4 (sense 42-mer, exactly complementary to primer 4CT) and oligo G $\alpha$ <sub>q</sub>i58R, (antisense 21-mer, complementary to G $\alpha$ <sub>q</sub>i5 cDNA; sequence 5'-ctcgatctcgtcgttgatccg-3'). The two PCR products generated from the above two reactions were annealed together in equimolar ratios in the presence of the external primers mG4-2028R and G $\alpha$ <sub>q</sub>i58R, and Pfu DNA polymerase (Stratagene).

The resulting chimeric PCR product was digested sequentially with KpnI and NotI (New England Biolabs) and subcloned into ph8SPmGluR4 prepared with the same two restriction enzymes. This intermediate construct was known as ph8SPmGluR4//CaR(no stop). In the final cloning step, a fragment containing the G $\alpha$ <sub>q</sub>i5 cDNA was released from phmGluR8//CaR\*AAA\*G $\alpha$ <sub>q</sub>i5 using the restriction enzymes ApaI and NotI (both New England Biolabs) and subcloned into the ph8SPmGluR4//CaR(no stop) construct, which was prepared with the same restriction enzymes. The sequence of the resultant chimeric construct, ph8SPmGluR4//CaR\*AAA\*G $\alpha$ <sub>q</sub>i5, was verified by ABI automated DNA sequence analysis.

#### VIII. phmGluR8//CaR Construct

This chimera contains the extracellular and transmembrane domains of human mGluR8 linked to the intracellular cytoplasmic tail domain of the human CaR. The

chimeric junction between hmGluR8 and the CaR was created using a recombinant PCR strategy similar to those described above.

The first reaction used two primers, CH5A (sense 19-mer, corresponding to nucleotides 2187-2205 of human mGluR8, Stormann *et al.*, U.S. Patent Nos. 6,051,688, 6,077,675, and 6,084,084) and the hybrid primer CH5B (antisense 40-mer, containing 22 nucleotides complementary to nucleotides 2523 – 2544 of human mGluR8, and 18 nucleotides complementary to nucleotides 2602-2619 of the human CaR). These primers were used to amplify a 375 bp PCR fragment of human mGluR8. In a separate PCR reaction using phCaR in the BlueScript SK(-) plasmid as template, a 750 bp fragment of the human CaR was amplified using a hybrid primer CH5C (sense 40-mer, exactly complementary to primer CH5B) and the T3 primer commercially available from Stratagene.

The two PCR products generated from the above two reactions were annealed together in equimolar ratios in the presence of the external primers CH5A and T3, and the Pfu DNA polymerase (Stratagene). The resulting chimeric PCR product was digested with BsrG I and Xba I (New England Biolabs) and subcloned into pmGluR8 digested with the same two restriction enzymes. The sequence of the resultant chimeric construct, pmGluR8//CaR, was verified by DNA sequence analysis.

#### IX. mGluR8//CaR\*Gα<sub>q</sub>i5 Construct

This construct contains the hmGluR8//CaR chimeric receptor fused to human Gα<sub>q</sub>i5. The chimeric junction between the C-terminus of hmGluR8//CaR and the N-terminus of Gα<sub>q</sub>i5 was created using a recombinant PCR strategy similar to that described above for the construction of phmGluR2//CaR\*Gα<sub>q</sub>i5.

The first reaction used two primers, CRP10A (sense 18-mer, corresponding to nucleotides 2812-2829 of phCaR) and the hybrid primer Gqi5/CaR (antisense 40-mer, containing 21 nucleotides complementary to nucleotides 3214-3234 phCaR, and 19 nucleotides complementary to nucleotides 1-19 of pGα<sub>q</sub>i5). These primers were used to amplify a 441 bp PCR fragment of hmGluR8//CaR.

In a separate PCR reaction all of Gα<sub>q</sub>i5 was amplified using a hybrid primer CaR/Gqi5 (sense 40-mer, exactly complementary to primer Gqi5/CaR) and the Apa I-mut primer (20-mer). The two PCR products generated from the above two reactions were

annealed together in equimolar ratios in the presence of the external primers CRP10A and Apa I-mut, and the Pfu DNA polymerase (Stratagene).

The resulting chimeric PCR product was digested with BstE II and Apa I (New England Biolabs) and subcloned into pmGluR8//CaR digested with the same two  
 5 restriction enzymes. The sequence of the resultant chimeric fusion construct, pmGluR8//CaR\*Gα<sub>qi5</sub>, was verified by DNA sequence analysis.

#### Example 2: Functional Expression of CaR/GABA<sub>B</sub>R2

In vitro transcribed RNA (7 ng) encoding a chimeric CaR/GABA<sub>B</sub>R2 (CaR  
 10 extracellular and transmembrane domains, and intracellular GABA<sub>B</sub>R2 domain) was co-injected with *in vitro* transcribed RNA (2 ng) encoding G 15 into *Xenopus* oocytes. Following a 72-hour incubation, the oocytes were voltage-clamped using standard electrophysiological techniques (Hille, B., Ionic Channels of Excitable Membranes, pp.30-33, Sinauer Associates, Inc., Sunderland, Ma., 1992). Activation of the chimeric  
 15 receptor was detected by increases in the calcium-activated chloride current.

Application of the CaR activator 100 Gd<sup>3+</sup>, resulted in reversible, oscillatory increases in the calcium-activated chloride current as shown in Figure 8. These data demonstrate the functional response of the chimeric CaR/GABA<sub>B</sub>R2 receptor upon activation via a site within the CaR extracellular domain. In this assay, the G 15 subunit  
 20 acts to promote signal transduction through intracellular pathways that mobilize intracellular Ca<sup>++</sup>.

#### Example 3: Expression of Different G-Protein Fusion Receptors

The ability of different G-protein fusions to transduce signal resulting from ligand  
 25 binding is shown in Figure 15. The different G-protein fusion receptors used in this example were as follows: mGluR2//CaR\*Gqi5 (SEQ. ID. NO. 37), CaR/mGluR2\*Gqi5 (SEQ. ID. NO. 33), and mGluR8//CaR\*Gqi5 SEQ. ID. NO. 41.

Oocytes suitable for injection were obtained from adult female *Xenopus laevis* toads using procedures described in C. J. Marcus-Sekura and M. J. M. Hitchcock,  
 30 Methods in Enzymology, Vol. 152 (1987).

Receptor fusion cRNAs were dissolved in water and 50 nl (12.5 ng/oocyte) were injected into individual oocytes. Following injection, oocytes were incubated at 16°C in MBS containing 1 mM CaCl<sub>2</sub> for 2 to 7 days prior to electrophysiological recording.

Test substances were applied by superfusion at a flow rate of about 5 ml/min. Receptor fusion activation was determined by measuring the increase in calcium-activated chloride current ( $I_{Cl}$ ). Increases in  $I_{Cl}$  were quantified by measuring the peak inward current stimulated by activating agent, relative to the holding current at -60 mV.

- 5 Application of 100  $\mu$ M L-glutamate elicited a response from the mGluR2//CaR\*G $\alpha$ q5 and mGluR8//CaR\*G $\alpha$ q5. Application of 100  $\mu$ M Gd<sup>3+</sup> activated the CaR/mGluR2\*Gq5.

#### Example 4: Expression of Different G-Protein Fusion Receptors in Mammalian Cells

10

HEK293 cells were transiently transfected with the p8SPhmGluR4//CaR\*AAA\*G $\alpha$ q5 or phmGluR8//CaR\*G $\alpha$ q5 plasmid DNAs using the following protocol. Initially, 150 cm<sup>2</sup> tissue culture flasks containing HEK293 cells at 75% confluence were transfected with 24  $\mu$ g of plasmid DNA using Gibco BRL Life Technologies' Lipofectamine reagent. Following liposomal gene delivery the cells were allowed to recover for 24 hours. They were then plated overnight at 100,000 cells per well in black, clear bottom, Collagen I coated 96-well plates (Becton Dickinson, Biocoat) using DMEM supplemented with 10% fetal bovine serum (Hyclone Laboratories). The cells were assayed for function 48 hours after transient transfection.

15

20

On the day of the assay, tissue culture medium was aspirated from the wells of a 96-well plate and 80  $\mu$ L of Assay Buffer (Assay Buffer is: 20 mM HEPES, 146 mM NaCl, 5 mM KCl, 1 mM MgCl<sub>2</sub>, 1 mM CaCl<sub>2</sub>, 1 mg/ml BSA, 1 mg/ml glucose, pH 7.4) supplemented with 6  $\mu$ M of the Ca<sup>2+</sup>-sensitive dye, Fluo-3 AM (Molecular Probes) and 0.025% Pluronic (Molecular Probes) was added to each well.

25

The plate was then incubated in the dark for 1 hour at room temperature to efficiently load the cells with Fluo-3. At the end of the incubation, extracellular Fluo-3 was removed by washing the plate with Assay Buffer. Assay Buffer was added back to each well (final volume = 160  $\mu$ L) prior to beginning the assay. The plate was loaded into a fluorescence imaging plate reader (FLIPR) robotic device (Molecular Devices) with the laser setting at 0.8 Watts. At a time of 15 seconds after initiation of the assay, 40  $\mu$ L of Assay Buffer containing 150  $\mu$ M L-AP4 was added to the 160  $\mu$ L of Assay Buffer in each well of the plate to yield a final concentration of 30  $\mu$ M L-AP4.

30



Relative fluorescence intensity (excitation  $\lambda = 488$  nm / emission  $\lambda = 510$  nm) was monitored at relevant time intervals throughout the assay period to measure L-AP4-induced receptor activation.

- 5 Other embodiments are within the following claims. Thus, while several embodiments have been shown and described, various modifications may be made, without departing from the spirit and scope of the present invention.

11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47 48 49 50 51 52 53 54 55 56 57 58 59 60 61 62 63 64 65 66 67 68 69 70 71 72 73 74 75 76 77 78 79 80 81 82 83 84 85 86 87 88 89 90 91 92 93 94 95 96 97 98 99 100 101 102 103 104 105 106 107 108 109 110 111 112 113 114 115 116 117 118 119 120 121 122 123 124 125 126 127 128 129 130 131 132 133 134 135 136 137 138 139 140 141 142 143 144 145 146 147 148 149 150 151 152 153 154 155 156 157 158 159 160 161 162 163 164 165 166 167 168 169 170 171 172 173 174 175 176 177 178 179 180 181 182 183 184 185 186 187 188 189 190 191 192 193 194 195 196 197 198 199 200 201 202 203 204 205 206 207 208 209 210 211 212 213 214 215 216 217 218 219 220 221 222 223 224 225 226 227 228 229 230 231 232 233 234 235 236 237 238 239 240 241 242 243 244 245 246 247 248 249 250 251 252 253 254 255 256 257 258 259 260 261 262 263 264 265 266 267 268 269 270 271 272 273 274 275 276 277 278 279 280 281 282 283 284 285 286 287 288 289 290 291 292 293 294 295 296 297 298 299 300 301 302 303 304 305 306 307 308 309 310 311 312 313 314 315 316 317 318 319 320 321 322 323 324 325 326 327 328 329 330 331 332 333 334 335 336 337 338 339 340 341 342 343 344 345 346 347 348 349 350 351 352 353 354 355 356 357 358 359 360 361 362 363 364 365 366 367 368 369 370 371 372 373 374 375 376 377 378 379 380 381 382 383 384 385 386 387 388 389 390 391 392 393 394 395 396 397 398 399 400 401 402 403 404 405 406 407 408 409 410 411 412 413 414 415 416 417 418 419 420 421 422 423 424 425 426 427 428 429 430 431 432 433 434 435 436 437 438 439 440 441 442 443 444 445 446 447 448 449 450 451 452 453 454 455 456 457 458 459 460 461 462 463 464 465 466 467 468 469 470 471 472 473 474 475 476 477 478 479 480 481 482 483 484 485 486 487 488 489 490 491 492 493 494 495 496 497 498 499 500 501 502 503 504 505 506 507 508 509 510 511 512 513 514 515 516 517 518 519 520 521 522 523 524 525 526 527 528 529 530 531 532 533 534 535 536 537 538 539 540 541 542 543 544 545 546 547 548 549 550 551 552 553 554 555 556 557 558 559 560 561 562 563 564 565 566 567 568 569 570 571 572 573 574 575 576 577 578 579 580 581 582 583 584 585 586 587 588 589 590 591 592 593 594 595 596 597 598 599 600 601 602 603 604 605 606 607 608 609 610 611 612 613 614 615 616 617 618 619 620 621 622 623 624 625 626 627 628 629 630 631 632 633 634 635 636 637 638 639 640 641 642 643 644 645 646 647 648 649 650 651 652 653 654 655 656 657 658 659 660 661 662 663 664 665 666 667 668 669 670 671 672 673 674 675 676 677 678 679 680 681 682 683 684 685 686 687 688 689 690 691 692 693 694 695 696 697 698 699 700 701 702 703 704 705 706 707 708 709 710 711 712 713 714 715 716 717 718 719 720 721 722 723 724 725 726 727 728 729 730 731 732 733 734 735 736 737 738 739 740 741 742 743 744 745 746 747 748 749 750 751 752 753 754 755 756 757 758 759 760 761 762 763 764 765 766 767 768 769 770 771 772 773 774 775 776 777 778 779 780 781 782 783 784 785 786 787 788 789 790 791 792 793 794 795 796 797 798 799 800 801 802 803 804 805 806 807 808 809 810 811 812 813 814 815 816 817 818 819 820 821 822 823 824 825 826 827 828 829 830 831 832 833 834 835 836 837 838 839 840 841 842 843 844 845 846 847 848 849 850 851 852 853 854 855 856 857 858 859 860 861 862 863 864 865 866 867 868 869 870 871 872 873 874 875 876 877 878 879 880 881 882 883 884 885 886 887 888 889 890 891 892 893 894 895 896 897 898 899 900 901 902 903 904 905 906 907 908 909 910 911 912 913 914 915 916 917 918 919 920 921 922 923 924 925 926 927 928 929 930 931 932 933 934 935 936 937 938 939 940 941 942 943 944 945 946 947 948 949 950 951 952 953 954 955 956 957 958 959 960 961 962 963 964 965 966 967 968 969 970 971 972 973 974 975 976 977 978 979 980 981 982 983 984 985 986 987 988 989 990 991 992 993 994 995 996 997 998 999 1000 1001 1002 1003 1004 1005 1006 1007 1008 1009 1010 1011 1012 1013 1014 1015 1016 1017 1018 1019 1020 1021 1022 1023 1024 1025 1026 1027 1028 1029 1030 1031 1032 1033 1034 1035 1036 1037 1038 1039 1040 1041 1042 1043 1044 1045 1046 1047 1048 1049 1050 1051 1052 1053 1054 1055 1056 1057 1058 1059 1060 1061 1062 1063 1064 1065 1066 1067 1068 1069 1070 1071 1072 1073 1074 1075 1076 1077 1078 1079 1080 1081 1082 1083 1084 1085 1086 1087 1088 1089 1090 1091 1092 1093 1094 1095 1096 1097 1098 1099 1100 1101 1102 1103 1104 1105 1106 1107 1108 1109 1110 1111 1112 1113 1114 1115 1116 1117 1118 1119 1120 1121 1122 1123 1124 1125 1126 1127 1128 1129 1130 1131 1132 1133 1134 1135 1136 1137 1138 1139 1140 1141 1142 1143 1144 1145 1146 1147 1148 1149 1150 1151 1152 1153 1154 1155 1156 1157 1158 1159 1160 1161 1162 1163 1164 1165 1166 1167 1168 1169 1170 1171 1172 1173 1174 1175 1176 1177 1178 1179 1180 1181 1182 1183 1184 1185 1186 1187 1188 1189 1190 1191 1192 1193 1194 1195 1196 1197 1198 1199 1200 1201 1202 1203 1204 1205 1206 1207 1208 1209 1210 1211 1212 1213 1214 1215 1216 1217 1218 1219 1220 1221 1222 1223 1224 1225 1226 1227 1228 1229 1230 1231 1232 1233 1234 1235 1236 1237 1238 1239 1240 1241 1242 1243 1244 1245 1246 1247 1248 1249 1250 1251 1252 1253 1254 1255 1256 1257 1258 1259 1260 1261 1262 1263 1264 1265 1266 1267 1268 1269 1270 1271 1272 1273 1274 1275 1276 1277 1278 1279 1280 1281 1282 1283 1284 1285 1286 1287 1288 1289 1290 1291 1292 1293 1294 1295 1296 1297 1298 1299 1300 1301 1302 1303 1304 1305 1306 1307 1308 1309 1310 1311 1312 1313 1314 1315 1316 1317 1318 1319 1320 1321 1322 1323 1324 1325 1326 1327 1328 1329 1330 1331 1332 1333 1334 1335 1336 1337 1338 1339 1340 1341 1342 1343 1344 1345 1346 1347 1348 1349 1350 1351 1352 1353 1354 1355 1356 1357 1358 1359 1360 1361 1362 1363 1364 1365 1366 1367 1368 1369 1370 1371 1372 1373 1374 1375 1376 1377 1378 1379 1380 1381 1382 1383 1384 1385 1386 1387 1388 1389 1390 1391 1392 1393 1394 1395 1396 1397 1398 1399 1400 1401 1402 1403 1404 1405 1406 1407 1408 1409 1410 1411 1412 1413 1414 1415 1416 1417 1418 1419 1420 1421 1422 1423 1424 1425 1426 1427 1428 1429 1430 1431 1432 1433 1434 1435 1436 1437 1438 1439 1440 1441 1442 1443 1444 1445 1446 1447 1448 1449 1450 1451 1452 1453 1454 1455 1456 1457 1458 1459 1460 1461 1462 1463 1464 1465 1466 1467 1468 1469 1470 1471 1472 1473 1474 1475 1476 1477 1478 1479 1480 1481 1482 1483 1484 1485 1486 1487 1488 1489 1490 1491 1492 1493 1494 1495 1496 1497 1498 1499 1500 1501 1502 1503 1504 1505 1506 1507 1508 1509 1510 1511 1512 1513 1514 1515 1516 1517 1518 1519 1520 1521 1522 1523 1524 1525 1526 1527 1528 1529 1530 1531 1532 1533 1534 1535 1536 1537 1538 1539 1540 1541 1542 1543 1544 1545 1546 1547 1548 1549 1550 1551 1552 1553 1554 1555 1556 1557 1558 1559 1560 1561 1562 1563 1564 1565 1566 1567 1568 1569 1570 1571 1572 1573 1574 1575 1576 1577 1578 1579 1580 1581 1582 1583 1584 1585 1586 1587 1588 1589 1590 1591 1592 1593 1594 1595 1596 1597 1598 1599 1600 1601 1602 1603 1604 1605 1606 1607 1608 1609 1610 1611 1612 1613 1614 1615 1616 1617 1618 1619 1620 1621 1622 1623 1624 1625 1626 1627 1628 1629 1630 1631 1632 1633 1634 1635 1636 1637 1638 1639 1640 1641 1642 1643 1644 1645 1646 1647 1648 1649 1650 1651 1652 1653 1654 1655 1656 1657 1658 1659 1660 1661 1662 1663 1664 1665 1666 1667 1668 1669 1670 1671 1672 1673 1674 1675 1676 1677 1678 1679 1680 1681 1682 1683 1684 1685 1686 1687 1688 1689 1690 1691 1692 1693 1694 1695 1696 1697 1698 1699 1700 1701 1702 1703 1704 1705 1706 1707 1708 1709 1710 1711 1712 1713 1714 1715 1716 1717 1718 1719 1720 1721 1722 1723 1724 1725 1726 1727 1728 1729 1730 1731 1732 1733 1734 1735 1736 1737 1738 1739 1740 1741 1742 1743 1744 1745 1746 1747 1748 1749 1750 1751 1752 1753 1754 1755 1756 1757 1758 1759 1760 1761 1762 1763 1764 1765 1766 1767 1768 1769 1770 1771 1772 1773 1774 1775 1776 1777 1778 1779 1780 1781 1782 1783 1784 1785 1786 1787 1788 1789 1790 1791 1792 1793 1794 1795 1796 1797 1798 1799 1800 1801 1802 1803 1804 1805 1806 1807 1808 1809 1810 1811 1812 1813 1814 1815 1816 1817 1818 1819 1820 1821 1822 1823 1824 1825 1826 1827 1828 1829 1830 1831 1832 1833 1834 1835 1836 1837 1838 1839 1840 1841 1842 1843 1844 1845 1846 1847 1848 1849 1850 1851 1852 1853 1854 1855 1856 1857 1858 1859 1860 1861 1862 1863 1864 1865 1866 1867 1868 1869 1870 1871 1872 1873 1874 1875 1876 1877 1878 1879 1880 1881 1882 1883 1884 1885 1886 1887 1888 1889 1890 1891 1892 1893 1894 1895 1896 1897 1898 1899 1900 1901 1902 1903 1904 1905 1906 1907 1908 1909 1910 1911 1912 1913 1914 1915 1916 1917 1918 1919 1920 1921 1922 1923 1924 1925 1926 1927 1928 1929 1930 1931 1932 1933 1934 1935 1936 1937 1938 1939 1940 1941 1942 1943 1944 1945 1946 1947 1948 1949 1950 1951 1952 1953 1954 1955 1956 1957 1958 1959 1960 1961 1962 1963 1964 1965 1966 1967 1968 1969 1970 1971 1972 1973 1974 1975 1976 1977 1978 1979 1980 1981 1982 1983 1984 1985 1986 1987 1988 1989 1990 1991 1992 1993 1994 1995 1996 1997 1998 1999 2000 2001 2002 2003 2004 2005 2006 2007 2008 2009 2010 2011 2012 2013 2014 2015 2016 2017 2018 2019 2020 2021 2022 2023 2024 2025 2026 2027 2028 2029 2030 2031 2032 2033 2034 2035 2036 2037 2038 2039 2040 2041 2042 2043 2044 2045 2046 2047 2048 2049 2050 2051 2052 2053 2054 2055 2056 2057 2058 2059 2060 2061 2062 2063 2064 2065 2066 2067 2068 2069 2070 2071 2072 2073 2074 2075 2076 2077 2078 2079 2080 2081 2082 2083 2084 2085 2086 2087 2088 2089 2090 2091 2092 2093 2094 2095 2096 2097 2098 2099 2100 2101 2102 2103 2104 2105 2106 2107 2108 2109 2110 2111 2112 2113 2114 2115 2116 2117 2118 2119 2120 2121 2122 2123 2124 2125 2126 2127 2128 2129 2130 2131 2132 2133 2134 2135 2136 2137 2138 2139 2140 2141 2142 2143 2144 2145 2146 2147 2148 2149 2150 2151 2152 2153 2154 2155 2156 2157 2158 2159 2160 2161 2162 2163 2164 2165 2166 2167 2168 2169 2170 2171 2172 2173 2174 2175 2176 2177 2178 2179 2180 2181 2182 2183 2184 2185 2186 2187 2188 2189 2190 2191 2192 2193 2194 2195 2196 2197 2198 2199 2200 2201 2202 2203 2204 2205 2206 2207 2208 2209 2210 2211 2212 2213 2214 2215 2216 2217 2218 2219 2220 2221 2222 2223 2224 2225 2226 2227 2228 2229 2230 2231 2232 2233 2234 2235 2236 2237 2238 2239 2240 2241 2242 2243 2244 2245 2246 2247 2248 2249 2250 2251 2252 2253 2254 2255 2256 2257 2258 2259 2260 2261 2262 2263 2264 2265 2266 2267 2268 2269 2270 2271 2272 2273 2274 2275 2276 2277 2278 2279 2280 2281 2282 2283 2284 2285 2286 2287 2288 2289 2290 2291 2292 2293 2294 2295 2296 2297 2298 2299 2300 2301 2302 2303 2304 2305 2306 2307 2308 2309 2310 2311 2312 2313 2314 2315 2316 2317 2318 2319 2320 2321 2322 2323 2324 2325 2326 2327 2328 2329 2330 2331 2332 2333 2334 2335 2336 2337 2338 2339 2340 2341 2342 2343 2344 2345 2346 2347 2348 2349 2350 2351 2352 2353 2354 2355 2356 2357 2358 2359 2360 2361 2362 2363 2364 2365 2366 2367 2368 2369 2370 2371 2372 2373 2374 2375 2376 2377 2378 2379 2380 2381 2382 2383 2384 2385 2386 2387 2388 2389 2390 2391 2392 2393 2394 2395 2396 2397 2398 2399 2400 2401 2402 2403 2404 2405 2406 2407 2408 2409 2410 2411 2412 2413 2414 2415 2416 2417 2418 2419 2420 2421 2422 2423 2424 2425 2426 2427 2428 2429 2430 2431 2432 2433 2434 2435 2436 2437 2438 2439 2440 2441 2442 2443 2444 2445 2446 2447 2448 2449 2450 2451 2452 2453 2454 2455 2456 2457 2458 2459 2460 2461 2462 2463 2464 2465 2466 2467 2468 2469 2470 2471 2472 2473 2474 2475 2476 2477 2478 2479 2480 2481 2482 2483 2484 2485 2486 2487 2488 2489 2490 2491 2492 2493 2494 2495 2496 2497 2498 2499 2500 2501 2502 2503 2504 2505 2506 2507 2508 2509 2510 2511 2512 2513 2514 2515 2516 2517 2518 2519 2520 2521 2522 2523 2524 2525 2526 2527 2528 2529 2530 2531 2532 2533 2534 2535 2536 2537 2538 2539 2540 2541 2542 2543 2544 2545 2546 2547 2548 2549 2550 2551 2552 2553 2554 2555 2556 2557 2558 2559 2560 2561 2562 2563 2564 2565 2566 2567 2568 2569 2570 2571 2572 2573 2574 2575 2576 2577 2578 2579 2580 2581 2582 2583 2584 2585 2586 2587 2588 2589 2590 2591 2592 2593 2594 2595 2596 2597 2598 2599 2600 2601 2602 2603 2604 2605 2606 2607 2608 2609 2610 2611 2612 2613 2614 2615 2616 2617 2618 2619 2620 2621 2622 2623 2624 2625 2626 2627 2628 2629 2630 2631 2632 2633 2634 2635 2636 2637 2638 2639 2640 2641 2642 2643 2644 2645 2646 2647 2648 2649 2650 2651 2

### Claims

1. A G-protein fusion receptor comprising

an extracellular domain comprising an extracellular domain amino acid sequence

substantially similar to either an extracellular CaR amino acid sequence, an extracellular mGluR amino acid sequence, or an extracellular GABA<sub>B</sub> receptor amino acid sequence;

a transmembrane domain joined to the carboxy terminus of said extracellular domain, said transmembrane domain comprising a transmembrane domain amino acid sequence substantially similar to either a transmembrane CaR amino acid sequence, a transmembrane mGluR amino acid sequence, or a transmembrane GABA<sub>B</sub> receptor amino acid sequence;

an intracellular domain joined to the carboxy terminus of said transmembrane domain comprising all or a portion of an intracellular amino acid sequence substantially similar to either an intracellular CaR amino acid sequence, an intracellular mGluR amino acid sequence, or an intracellular GABA<sub>B</sub> receptor amino acid sequence, provided that said portion is at least about 10 amino acids;

an optionally present linker joined to the carboxy terminus of said intracellular domain; and

a G-protein joined either to said intracellular domain or to said optionally present linker, provided that said G-protein is joined to said optionally present linker when said optionally present linker is present.

2. The G-protein fusion receptor of claim 1, wherein said extracellular domain consists of said extracellular domain amino acid sequence, said transmembrane domain consists of said transmembrane domain amino acid sequence; and said intracellular domain consists of said transmembrane domain amino acid sequence.

3. The G-protein fusion receptor of claim 2, wherein said optionally present linker is present and is a polypeptide 3 to 30 amino acids in length.

4. The G-protein fusion receptor of claim 2, wherein said optionally present linker is not present.

5. 1. The G-protein fusion receptor of claim 3 or 4, wherein said G-protein is selected from the group consisting of: G<sub>15</sub>, G<sub>16</sub>, Gqo5, and Gqi5

6. The G-protein fusion of claim 5, wherein any of said CaR sequence present is a human CaR sequence, any of said mGluR sequence present is from a human mGluR, and any of said GABA<sub>B</sub> receptor sequence present is from human mGluR.

7. A nucleic acid comprising a nucleotide sequence encoding for the G-protein fusion of any one of claims 1-6.

8. An expression vector comprising a nucleotide sequence encoding for the G-protein fusion of any one of claims 1-6 transcriptionally coupled to a promoter.

9. A recombinant cell comprising the expression vector of claim 8 and a cell wherein the G-protein fusion is expressed and is functional.

10. A recombinant cell produced by combining a vector comprising the nucleic acid of claim 9 and elements for introducing heterologous nucleic acid into a cell wherein the G-protein fusion receptor is expressed, and said cell.

11. A process for the production of a G-protein fusion receptor comprising: growing procaryotic or eukaryotic host cells comprising a nucleic acid sequence expressing the G-protein fusion receptor of any one of claims 1-6, under suitable nutrient conditions allowing for cell growth.

12. A method of measuring the ability of a compound to effect G-protein fusion activity comprising the steps of:

a) providing said compound to a cell expressing the G-protein fusion receptor of any one of claims 1-6, and

b) measuring the ability of said compound to affect the activity of said receptor as an indication of the ability of said compound to effect G-protein fusion receptor activity.

13. A chimeric receptor comprising

an extracellular domain comprising an extracellular domain amino acid sequence substantially similar to a sequence selected from the group consisting of SEQ ID NO: 1, SEQ ID NO: 2, SEQ ID NO: 3, SEQ ID NO: 4 and SEQ ID NO: 5;

a transmembrane domain comprising a transmembrane domain amino acid sequence substantially similar to a sequence selected from the group consisting of SEQ ID NO: 6, SEQ ID NO: 7, SEQ ID NO: 8, SEQ ID NO: 9, and SEQ ID NO: 10; and

an intracellular cytoplasmic domain comprising an intracellular domain amino acid sequence substantially similar to a sequence selected from the group consisting of SEQ ID NO: 11, SEQ ID NO: 12, SEQ ID NO: 13, and SEQ ID NO: 14;

wherein at least one domain is present which comprises an amino acid sequence substantially similar to a sequence selected from the group consisting of: SEQ ID NO: 2, SEQ ID NO: 3, SEQ ID NO: 4, SEQ ID NO: 7, SEQ ID NO: 8, and SEQ ID NO: 9, SEQ ID NO: 12, SEQ ID NO: 13, and SEQ ID NO: 14; and at least one domain is present which comprises an amino acid sequence substantially similar to a sequence selected from the group consisting of: SEQ ID NO: 1, SEQ ID NO: 5, SEQ ID NO: 6, SEQ ID NO: 10, SEQ ID NO: 11, and SEQ ID NO: 15.

14. The chimeric receptor of claim 13 wherein said extracellular domain has a sequence similarity of at least 90% with an amino acid sequence selected from the group consisting of SEQ ID NOs: 2, 3, and 4; said transmembrane domain has a sequence similarity of at least 90% with an amino acid sequence selected from the group consisting of SEQ ID Nos: 6, 7, 8, 9, and 10; and said intracellular domain has a sequence similarity of at least 90% with an amino acid sequence selected from the group consisting of SEQ ID NOs: 11 and 15.

15. The chimeric receptor of claim 14, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 2; said transmembrane domain has a sequence similarity of at least 90% with the amino acid sequence SEQ ID NO: 7; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 11.

16. The chimeric receptor of claim 14, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 3; said

transmembrane domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 8; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO 11.

5           17.     The chimeric receptor of claim 14, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence SEQ ID NO: 4; said transmembrane domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 9; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO 11.

10           18.     The chimeric receptor of claim 13, wherein said extracellular domain has a sequence similarity of at least 90% with an amino acid sequence selected from the group consisting of SEQ ID NOs: 1, 2, 3, 4 and 5; said transmembrane domain has a sequence similarity of at least 90% with an amino acid sequence selected from the group consisting of SEQ ID Nos: 7, 8, and 9; and said intracellular domain has a sequence similarity of at least 90% with an amino acid sequence selected from the group consisting of SEQ ID NOs: 11, 12, 13, 14, and 15.

15           19.     The chimeric receptor of claim 18, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 1; said transmembrane domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 7; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 11.

20           20.     The chimeric receptor of claim 18, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 1; said transmembrane domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 8; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO 11.

25           21.     The chimeric receptor of claim 18, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 1; said transmembrane domain has a sequence similarity of at least 90% with the amino acid

30

sequence of SEQ ID NO: 9; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO 11.

22. The chimeric receptor of claim 13, wherein said extracellular domain has a sequence similarity of at least 90% with an amino acid sequence selected from the group consisting of SEQ ID NOs: 1, 2, 3, 4, and 5; said transmembrane domain has a sequence similarity of at least 90% with an amino acid sequence selected from the group consisting of SEQ ID Nos: 6, 7, 8, 9, and 10; and said intracellular domain has a sequence similarity of at least 90% with an amino acid sequence selected from the group consisting of SEQ ID NOs: 12, 13, and 14.

23. The chimeric receptor of claim 22, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 1; said transmembrane domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 6; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 12.

24. The chimeric receptor of claim 22, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 1; said transmembrane domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 7; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 12.

25. The chimeric receptor of claim 22, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 1; said transmembrane domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 8; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 13.

26. The chimeric receptor of claim 22, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 1; said transmembrane domain has a sequence similarity of at least 90% with the amino acid

sequence of SEQ ID NO: 6; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 13.

27. The chimeric receptor of claim 22, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 1; said transmembrane domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 9; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 14.

28. The chimeric receptor of claim 22, wherein said extracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 1; said transmembrane domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 6; and said intracellular domain has a sequence similarity of at least 90% with the amino acid sequence of SEQ ID NO: 14.

29. The chimeric receptor of any one of claims 13-28, wherein said receptor functional couples to a G-protein.

30. The chimeric receptor of any one of claims 13-28, wherein said chimeric receptor consists of said extracellular domain, said transmembrane domain, said intracellular domain, and an optionally present G-protein  $\alpha$  subunit covalently joined to said intracellular domain.

31. The chimeric receptor of claim 30, wherein said chimeric receptor consists of said extracellular domain, said transmembrane domain, and said intracellular domain.

32. The chimeric receptor of claim 30, wherein said G-protein  $\alpha$  subunit consists of the amino acid sequence of SEQ ID Nos: 16 or 17.

33. A nucleic acid comprising a nucleotide sequence encoding for the chimeric receptor of any one of claims 13-32.

34. An expression vector comprising a nucleotide sequence encoding for the chimeric receptor of any one of claims 13-32 transcriptionally coupled to a promoter.

35. A recombinant cell comprising the expression vector of claim 34 and a cell wherein the chimeric receptor is expressed and is functional.

36. A recombinant cell produced by combining a vector comprising the nucleic acid of claim 33 and elements for introducing heterologous nucleic acid into a cell wherein the chimeric receptor is expressed, and said cell.

37. A process for the production of a chimeric receptor comprising:  
growing procaryotic or eukaryotic host cells comprising a nucleic acid sequence expressing the chimeric receptor of any one of claims 13-32, under suitable nutrient conditions allowing for cell growth.

38. A method of measuring the ability of a compound to effect GABA<sub>B</sub>R or mGluR activity comprising the steps of:

a) providing said compound to a cell expressing the chimeric receptor of any one of claims 13-32, and

b) measuring the ability of said compound to affect the activity of said receptor as an indication of the ability of said compound to effect GABA<sub>B</sub>R or mGluR activity.

39. The method of claim 38, wherein said method measures activity at a GABA<sub>B</sub>R.

40. The method of claim 38, wherein said method measures activity at a mGluR.

41. A fusion receptor polypeptide comprising a receptor and a G-protein  $\alpha$  subunit, wherein said G-protein  $\alpha$  subunit is fused to the intracellular domain of said receptor, provided that said receptor is not an adrenoreceptor.



## ABSTRACT

The present invention features G-protein fusion receptors and chimeric GABA<sub>B</sub> receptors (GABA<sub>B</sub>Rs), nucleic acid encoding such receptors, and the use of such receptors and nucleic acid. G-protein fusion receptors comprise at least one domain from a CaR, a mGluR, and/or a GABA<sub>B</sub> receptor fused directly or through a linker to a guanine nucleotide-binding protein (G-protein). Chimeric GABA<sub>B</sub>Rs comprise at least one of a GABA<sub>B</sub>R extracellular domain, a GABA<sub>B</sub>R transmembrane domain, or a GABA<sub>B</sub>R intracellular domain and one or more domains from a mGluR subtype 8 (mGluR8) and/or a CaR.

# ClustalW Formatted Alignments

```

SEQ ID 1  MAFYSCCWVLLALTWHTSAYGPDQR
SEQ ID 2  MLLLLLLAPLFLRPPGAGGAQTPNA
SEQ ID 3  MGPGAPFARVGVWPLPLLVMAGVA
SEQ ID 4  MASPRSSGQPGXP PPPPPARLLL
SEQ ID 5  MVC E GKRSASCP CFFLLTAKFYWIL
  
```

```

SEQ ID 1  AQKKGDIILGGLFPIHFGVAAKDQD
SEQ ID 2  TSEGCQIIHPPWEGGI RYRGLTRDQ
SEQ ID 3  PVWASHSPHLPRPHSRVP PHPSSER
SEQ ID 4  LLLLPLLLPLAPGAWGWARGAPRPP
SEQ ID 5  TMMQRTHSQEYAH S IRVDGDIILGG
  
```

```

SEQ ID 1  LKSRPESVECI RYNFRGFRWLQAMI
SEQ ID 2  VKA INFLPVDYEIEYVCRGEREVVG
SEQ ID 3  RAVYIGALFPM SGGWPGGQACQPAV
SEQ ID 4  PSSPPLSIMGLMPLTKEVAKGSI GR
SEQ ID 5  LFPVHAKGERGVPCGELKKEKGIHR
  
```

```

SEQ ID 1  FAIEEINSSPALLPNLT LGYRIFD T
SEQ ID 2  PKVRKCLANGSWTDM DTPSRCVRIC
SEQ ID 3  EMAL EDVNSRRDILPDYELKLIHHD
SEQ ID 4  GVLPAVELAIEQIRNESLLRPYFLD
SEQ ID 5  LEAMLYAIDQINKDPDLLSNITLGV
  
```

```

SEQ ID 1  CNTVSKALEATLSFVAQNKIDSLNL
SEQ ID 2  SKSYLTLENGKVFLTGGDLPA LDGA
SEQ ID 3  SKCDPGQATKYLYELLYNDPIKII L
SEQ ID 4  LRLYDTECDNAKGLKAFYDAIKYGP
SEQ ID 5  RILDTC SRDTYALEQSLTFVQALIE
  
```

```

SEQ ID 1  DEFCNCSEHIPSTIAVVGATGSGVS
SEQ ID 2  RVDFRCDPDFHLVGSSRSIC SQGQW
SEQ ID 3  MPGCSSVSTLVAEAARMWN LIVLSY
SEQ ID 4  NHLMVFGGVCPSVTSIIAESLQGW N
SEQ ID 5  KDASDVKCAN G D PPIFTKPKI SGV
  
```

```

SEQ ID 1  TAVANLLGLFYIPQVSYASSSRLLS
SEQ ID 2  STPKPHCQVNRTPHSERRAVYIGAL
SEQ ID 3  GSSSPALSNRQRFPTFFRTHPSATL
SEQ ID 4  LVQLSFAATT PVLADKKKYPYFFRT
SEQ ID 5  IGAAASSVSIMVANILRLFKIPQIS
  
```

Figure 1a

SEQ ID 1 NKNQFKSFLRTIPNDEHQATAMADI  
 SEQ ID 2 FPMSSGGWPGGQACQPAVEMALEDVN  
 SEQ ID 3 HNPTRVKLF EKWGWKKIATIQQTTE  
 SEQ ID 4 VPSDNAVNPAI LKLLKH YQWKRVGT  
 SEQ ID 5 YASTAPELSDNTRYDFFSRVVPD S

SEQ ID 1 IEYFRWNWVG TIAADDDYGRPGIEK  
 SEQ ID 2 SRRDILPDYELKLIHHDSKCDPGQA  
 SEQ ID 3 VFTSTLDDLEERVKEAGIEITFRQS  
 SEQ ID 4 LTQDVQRFS EVRNDLTGVLYGEDIE  
 SEQ ID 5 YQAQAMVDIVTALGWNYVSTLASEG

SEQ ID 1 FREEABERDICI D FSELISQYSDEE  
 SEQ ID 2 TKYLYELL YNDPIKII LMPGCSSVS  
 SEQ ID 3 FFSDPAVPVKNLKRQDARIIVGLFY  
 SEQ ID 4 ISDTESFSNDPCTSVKKLKGNDVRI  
 SEQ ID 5 NYGESGV EAF TQISREIGGVCI AQS

SEQ ID 1 EIQHVVEVIQNSTAKVIVVFSSGPD  
 SEQ ID 2 TLVABAAARMWNLI VLSYSSSSPALS  
 SEQ ID 3 ETEARKVFCEVYKERLFGKKYVWFL  
 SEQ ID 4 ILGQFDQNMAAKVFCCAYEENMYGS  
 SEQ ID 5 QKIPREPRPGEF EKIIKRLLLETPNA

SEQ ID 1 LEPLIKEIVRRNITGKIWLASEAWA  
 SEQ ID 2 NRQRFPTFFRTHPSATLHNPTRVKL  
 SEQ ID 3 IGWYADNWFKIYDPSINCTVDEMTE  
 SEQ ID 4 KYQWII PGWYEP S WVEQVHTEANSS  
 SEQ ID 5 RAVIMFANEDDIRRI LEAAKKLNQS

SEQ ID 1 SSSLIAMPQYFHVVG GTIGFALKAG  
 SEQ ID 2 FEKWWGWKKIATIQQTTEVFTSTLDD  
 SEQ ID 3 AVEGHITTEIVMLNPANTRSI SNMT  
 SEQ ID 4 RCLRKNNLLAAMEGYIGVDFEPLSSK  
 SEQ ID 5 GHFLWIGSDSWGSKIAPVYQQEEIA

SEQ ID 1 QIPGFR EFLKKVHPRKSVHNGFAKE  
 SEQ ID 2 LEERVKEAGIEITFRQSFFSDPAVP  
 SEQ ID 3 SQEFVEKLT KRLKRHP EETGGFQEA  
 SEQ ID 4 QIKTISGKTPQQYEREYNNKRSGVG  
 SEQ ID 5 EGAVTILPKRASIDGFDRYFRSRTL

Figure 1b

SEQ ID 1 F W E E T F N C H L Q E G A K G P L P V D T F L R  
 SEQ ID 2 V K N L K R Q D A R I I V G L F Y E T E A R K V F  
 SEQ ID 3 P L A Y D A I W A L A L A L N K T S G G G G R S G  
 SEQ ID 4 P S K F H G Y A Y D G I W V I A K T L Q R A M E T  
 SEQ ID 5 A N N R R N V W F A E F W E E N F G C K L G S H G

SEQ ID 1 G H E E S G D R F S N S S T A F R P L C T G D E N  
 SEQ ID 2 C E V Y K E R L F G K K Y V W F L I G W Y A D N W  
 SEQ ID 3 V R L E D F N Y N N Q T I T D Q I Y R A M N S S S  
 SEQ ID 4 L H A S S R H Q R I Q D F N Y T D H T L G R I I L  
 SEQ ID 5 K R N S H I K K C T G L E R I A R D S S Y E Q E G

SEQ ID 1 I S S V E T P Y I D Y T H L R I S Y N V Y L A V Y  
 SEQ ID 2 F K I Y D P S I N C T V D E M T E A V E G H I T T  
 SEQ ID 3 F E G V S G H V V F D A S G S R M A W T L I E Q L  
 SEQ ID 4 N A M N E T N F F G V T G Q V V F R N G E R M G T  
 SEQ ID 5 K V Q F V I D A V Y S M A Y A L H N M H K D L C P

SEQ ID 1 S I A H A L Q D I Y T C L P G R G L F T N G S C A  
 SEQ ID 2 E I V M L N P A N T R S I S N M T S Q E F V E K L  
 SEQ ID 3 Q G G S Y K K I G Y Y D S T K D D L S W S K T D K  
 SEQ ID 4 I K F T Q F Q D S R E V K V G E Y N A V A D T L E  
 SEQ ID 5 G Y I G L C P R M S T I D G K E L L G Y I R A V N

SEQ ID 1 D I K K V E A W Q V L K H L R H L N F T N N M G B  
 SEQ ID 2 T K R L K R H P E E T G G F Q E A P L A Y D A I W  
 SEQ ID 3 W I G G S P P A D Q T L V I K T F R F L S Q K  
 SEQ ID 4 I I N D T I R F Q G S E P P K D K T I I L E Q L R  
 SEQ ID 5 F N G S A G T P V T F N E N G D A P G R Y D I F Q

SEQ ID 1 Q V T F D E C G D L V G N Y S I I N W H L S P E D  
 SEQ ID 2 A L A L A L N K T S G G G G R S G V R L E D F N Y  
 SEQ ID 3  
 SEQ ID 4 K I S L P  
 SEQ ID 5 Y Q I T N K S T E Y K V I G H W T N Q L H L K V E

SEQ ID 1 G S I V F K E V G Y Y N V Y A K K G E R L F I N E  
 SEQ ID 2 N N Q T I T D Q I Y R A M N S S S F E G V S G H V  
 SEQ ID 3  
 SEQ ID 4  
 SEQ ID 5 D M Q W A H R E H T H P A S V C S L P C K P G E R

Figure 1c

SEQ ID 1 E K I L W S G F S R E V P F S N C S R D C L A G T  
SEQ ID 2 V F D A S G S R M A W T L I E Q L Q G G S Y K K I  
SEQ ID 3  
SEQ ID 4  
SEQ ID 5 K K T V K G V P C C W H C E R C E G Y N Y Q V D E

SEQ ID 1 R K G I I E G E P T C C F E C V E C P D G E Y S D  
SEQ ID 2 G Y Y D S T K D D L S W S K T D K W I G G S P P A  
SEQ ID 3  
SEQ ID 4  
SEQ ID 5 L S C E L C P L D Q R P N M N R T G C Q L I P I I

SEQ ID 1 E T D A S A C N K C P D D F W S N E N H T S C I A  
SEQ ID 2 D Q T L V I K T F R F L S Q K  
SEQ ID 3  
SEQ ID 4  
SEQ ID 5 K L E W H S P W

SEQ ID 1 K E I E F L S W T E P F  
SEQ ID 2  
SEQ ID 3  
SEQ ID 4  
SEQ ID 5

Figure 1d

FIGURE 2a

SEQ ID 6 G I A L T L F A V L G I F L T A F V L G V F I K F R N T P I  
SEQ ID 7 L F I S V S V L S S L G I V L A V V C L S F N I Y N S H V R  
SEQ ID 8 L F I S V S V L S S L G I V L A V V C L S F N I Y N S H V R  
SEQ ID 9 L Y S I L S A L T I L G M I M A S A F L F F N I K N R N Q K  
SEQ ID 10 A V V P V F V A I L G I I A T T F V I V T F V R Y N D T P I

SEQ ID 6 V K A T N R E L S Y L L L F S L L C C F S S S L F F I G E P  
SEQ ID 7 Y I Q N S Q P N L N N L T A V G C S L A L A A V F P L G L D  
SEQ ID 8 Y I Q N S Q P N L N N L T A V G C S L A L A A V F P L G L D  
SEQ ID 9 L I K M S S P Y M N N L I I L G G M L S Y A S I F L F G L D  
SEQ ID 10 V R A S G R E L S Y V L L T G I F L C Y S I T F L M I A A P

SEQ ID 6 Q D W T C R L R Q P A F G I S F V L C I S C I L V K T N R V  
SEQ ID 7 G Y H I G R N Q F P F V C Q A R L W L L G L G F S L G Y G S  
SEQ ID 8 G Y H I G R N Q F P F V C Q A R L W L L G L G F S L G Y G S  
SEQ ID 9 G S F V S E K T F E T L C T V R T W I L T V G Y T T A F G A  
SEQ ID 10 D T I I C S F R R V F L G L G M C F S Y A A L L T K T N R I

SEQ ID 6 L L V F E A K I P T S F H R K W W G L N L Q F L L V F L C T  
SEQ ID 7 M F T K I W W V H T V F T K K E E K K E W R K T L E P W K L  
SEQ ID 8 M F T K I W W V H T V F T K K E E K K E W R K T L E P W K L  
SEQ ID 9 M F A K T W R V H A I F K N V K M K K K I I K D Q K L L V I  
SEQ ID 10 H R I F E Q G K K S V T A P K F I S P A S Q L V I T F S L I

SEQ ID 6 F M Q I V I C V I W L Y T A P P S S Y R N Q E L E D E I I F  
SEQ ID 7 Y A T V G L L V G M D V L T L A I W Q I V D P L H R T I E T  
SEQ ID 8 Y A T V G L L V G M D V L T L A I W Q I V D P L H R T I E T  
SEQ ID 9 V G G M L L I D L C I L I C W Q A V D P L R R T V E K Y S M  
SEQ ID 10 S V Q L L G V F V W F V V D P P H I I I D Y G E Q R T L D P

SEQ ID 6 I T C H E G S L M A L G F L I G Y T C L L A A I C F F F A F  
SEQ ID 7 F A K E E P K E D I D V S I L P Q L E H C S S R K M N T W L  
SEQ ID 8 F A K E E P K E D I D V S I L P Q L E H C S S R K M N T W L  
SEQ ID 9 E P D P A G R D I S I R P L L E H C E N T H M T I W L G I V  
SEQ ID 10 E K A R G V L K C D I S D L S L I C S L G Y S I L L M V T C

SEQ ID 6 K S R K L P E N F N E A K F I T F S M L I F F I V W I S F I  
SEQ ID 7 G I F Y G Y K G L L L L L G I F L A Y E T K S V S T E K I N  
SEQ ID 8 G I F Y G Y K G L L L L L G I F L A Y E T K S V S T E K I N  
SEQ ID 9 Y A Y K G L L M L F G C F L A W E T R N V S I P A L N D S K  
SEQ ID 10 T V Y A I K T R G V P E T F N E A K P I G F T M Y T T C I I

[illegible]

```

SEQ ID 6      F F N K I Y I I L F
SEQ ID 7      D A A F A F A S L A I V F S S Y I T L V V L F V P K M
SEQ ID 8      D A A F A F A S L A I V F S S Y I T L V V L F V P K M
SEQ ID 9      F C I V A L V I I F C S T I T L C L V F V P K L
SEQ ID 10     L S A S V S L G M L Y M P K V Y I I I F

```

FIGURE 3a

SEQ ID 11 K P S R N T I E E V R C S T A A H A F K V A A R A T L R R S  
SEQ ID 12 R R L I T R G E W Q S E A Q D T M K T G S S T N N N E E E K  
SEQ ID 13 R R L I T R G E W Q S E A Q D T M K T G S S T N N N E E E K  
SEQ ID 14 I T L R T N P D A A T Q N R R F Q F T Q N Q K K E D S K T S  
SEQ ID 15 H P E Q N V Q K R K R S F K A V V T A A T M Q S K L I Q K G

SEQ ID 11 N V S R K R S S S L G G S T G S T P S S S I S S K S N S E D  
SEQ ID 12 S R L L E K E N R E L E K I I A E K E E R V S E L R H Q L Q  
SEQ ID 13 S R L L E K E N R E L E K I I A E K E E R V S E L R H Q L Q  
SEQ ID 14 T S V T S V N Q A S T S R L E G L Q S E N H R L R M K I T E  
SEQ ID 15 N D R P N G E V K S E L C E S L E T N S K S S V E F P M V K

SEQ ID 11 P F P Q P E R Q K Q Q Q P L A L T Q Q E Q Q Q Q P L T L P Q  
SEQ ID 12 S R Q Q L R S R R H P P T P P E P S G G L P R G P P E P P D  
SEQ ID 13 S R Q Q L R S R R H P P T P P E P S G G L P R G P P E P P D  
SEQ ID 14 L D K D L E E V T M Q L Q D T P E K T T Y I K Q N H Y Q E L  
SEQ ID 15 S G S T S

SEQ ID 11 Q Q R S Q Q Q P R C K Q K V I F G S G T V T F S L S F D E P  
SEQ ID 12 R L S C D G S R V H L L Y K  
SEQ ID 13 R L S C D G S R V H L L Y K  
SEQ ID 14 N D I L N L G N F T E S T D G G K A I L K N H L D Q N P Q L  
SEQ ID 15

SEQ ID 11 Q K N A M A H G N S T H Q N S L E A Q K S S D T L T R H Q P  
SEQ ID 12  
SEQ ID 13  
SEQ ID 14 Q W N T T E P S R T C K D P I E D I N S P E H I Q R R L S L  
SEQ ID 15

SEQ ID 11 L L P L Q C G E T D L D L T V Q E T G L Q G P V G G D Q R P  
SEQ ID 12  
SEQ ID 13  
SEQ ID 14 Q L P I L H H A Y L P S I G G V D A S C V S P C V S P T A S  
SEQ ID 15

SEQ ID 11 E V E D P E E L S P A L V V S S S Q S F V I S G G G S T V T  
SEQ ID 12  
SEQ ID 13  
SEQ ID 14 P R H R H V P P S F R V M V S G L  
SEQ ID 15



FIGURE 3'

SEQ ID 11 E N V V N S

SEQ ID 12

SEQ ID 13

SEQ ID 14

SEQ ID 15

11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47 48 49 50 51 52 53 54 55 56 57 58 59 60 61 62 63 64 65 66 67 68 69 70 71 72 73 74 75 76 77 78 79 80 81 82 83 84 85 86 87 88 89 90 91 92 93 94 95 96 97 98 99 100 101 102 103 104 105 106 107 108 109 110 111 112 113 114 115 116 117 118 119 120 121 122 123 124 125 126 127 128 129 130 131 132 133 134 135 136 137 138 139 140 141 142 143 144 145 146 147 148 149 150 151 152 153 154 155 156 157 158 159 160 161 162 163 164 165 166 167 168 169 170 171 172 173 174 175 176 177 178 179 180 181 182 183 184 185 186 187 188 189 190 191 192 193 194 195 196 197 198 199 200 201 202 203 204 205 206 207 208 209 210 211 212 213 214 215 216 217 218 219 220 221 222 223 224 225 226 227 228 229 230 231 232 233 234 235 236 237 238 239 240 241 242 243 244 245 246 247 248 249 250 251 252 253 254 255 256 257 258 259 260 261 262 263 264 265 266 267 268 269 270 271 272 273 274 275 276 277 278 279 280 281 282 283 284 285 286 287 288 289 290 291 292 293 294 295 296 297 298 299 300 301 302 303 304 305 306 307 308 309 310 311 312 313 314 315 316 317 318 319 320 321 322 323 324 325 326 327 328 329 330 331 332 333 334 335 336 337 338 339 340 341 342 343 344 345 346 347 348 349 350 351 352 353 354 355 356 357 358 359 360 361 362 363 364 365 366 367 368 369 370 371 372 373 374 375 376 377 378 379 380 381 382 383 384 385 386 387 388 389 390 391 392 393 394 395 396 397 398 399 400 401 402 403 404 405 406 407 408 409 410 411 412 413 414 415 416 417 418 419 420 421 422 423 424 425 426 427 428 429 430 431 432 433 434 435 436 437 438 439 440 441 442 443 444 445 446 447 448 449 450 451 452 453 454 455 456 457 458 459 460 461 462 463 464 465 466 467 468 469 470 471 472 473 474 475 476 477 478 479 480 481 482 483 484 485 486 487 488 489 490 491 492 493 494 495 496 497 498 499 500 501 502 503 504 505 506 507 508 509 510 511 512 513 514 515 516 517 518 519 520 521 522 523 524 525 526 527 528 529 530 531 532 533 534 535 536 537 538 539 540 541 542 543 544 545 546 547 548 549 550 551 552 553 554 555 556 557 558 559 560 561 562 563 564 565 566 567 568 569 570 571 572 573 574 575 576 577 578 579 580 581 582 583 584 585 586 587 588 589 590 591 592 593 594 595 596 597 598 599 600 601 602 603 604 605 606 607 608 609 610 611 612 613 614 615 616 617 618 619 620 621 622 623 624 625 626 627 628 629 630 631 632 633 634 635 636 637 638 639 640 641 642 643 644 645 646 647 648 649 650 651 652 653 654 655 656 657 658 659 660 661 662 663 664 665 666 667 668 669 670 671 672 673 674 675 676 677 678 679 680 681 682 683 684 685 686 687 688 689 690 691 692 693 694 695 696 697 698 699 700 701 702 703 704 705 706 707 708 709 710 711 712 713 714 715 716 717 718 719 720 721 722 723 724 725 726 727 728 729 730 731 732 733 734 735 736 737 738 739 740 741 742 743 744 745 746 747 748 749 750 751 752 753 754 755 756 757 758 759 760 761 762 763 764 765 766 767 768 769 770 771 772 773 774 775 776 777 778 779 780 781 782 783 784 785 786 787 788 789 790 791 792 793 794 795 796 797 798 799 800 801 802 803 804 805 806 807 808 809 810 811 812 813 814 815 816 817 818 819 820 821 822 823 824 825 826 827 828 829 830 831 832 833 834 835 836 837 838 839 840 841 842 843 844 845 846 847 848 849 850 851 852 853 854 855 856 857 858 859 860 861 862 863 864 865 866 867 868 869 870 871 872 873 874 875 876 877 878 879 880 881 882 883 884 885 886 887 888 889 890 891 892 893 894 895 896 897 898 899 900 901 902 903 904 905 906 907 908 909 910 911 912 913 914 915 916 917 918 919 920 921 922 923 924 925 926 927 928 929 930 931 932 933 934 935 936 937 938 939 940 941 942 943 944 945 946 947 948 949 950 951 952 953 954 955 956 957 958 959 960 961 962 963 964 965 966 967 968 969 970 971 972 973 974 975 976 977 978 979 980 981 982 983 984 985 986 987 988 989 990 991 992 993 994 995 996 997 998 999 1000

FIGURE 4a

SEQ. ID. NO. 16 M A R S L T W G C C P W C L T E E E K T A A R I D Q E I N R  
SEQ. ID. NO. 17 M A R S L T W R C C P W C L T E D E K A A A R V D Q E I N R

SEQ. ID. NO. 16 I L L E Q K K Q E R E E L K L L L L G P G E S G K S T F I K  
SEQ. ID. NO. 17 I L L E Q K K Q D R G E L K L L L L G P G E S G K S T F I K

SEQ. ID. NO. 16 Q M R I I H G V G Y S E E D R R A F R L L I Y Q N I F V S M  
SEQ. ID. NO. 17 Q M R I I H G A G Y S E E E R K G F R P L V Y Q N I F V S M

SEQ. ID. NO. 16 Q A M I D A M D R L Q I P F S R P D S K Q H A S L V M T Q D  
SEQ. ID. NO. 17 R A M I E A M E R L Q I P F S R P E S K H H A S L V M S Q D

SEQ. ID. NO. 16 P Y K V S T F E K P Y A V A M Q Y L W R D A G I R A C Y E R  
SEQ. ID. NO. 17 P Y K V T T F E K R Y A A A M Q W L W R D A G I R A C Y E R

SEQ. ID. NO. 16 R R E F H L L D S A V Y Y L S H L E R I S E D S Y I P T A Q  
SEQ. ID. NO. 17 R R E F H L L D S A V Y Y L S H L E R I T E E G Y V P T A Q

SEQ. ID. NO. 16 D V L R S R M P T T G I N E Y C F S V K K T K L R I V D V G  
SEQ. ID. NO. 17 D V L R S R M P T T G I N E Y C F S V Q K T N L R I V D V G

SEQ. ID. NO. 16 G Q R S E R R K W I H C F E N V I A L I Y L A S L S E Y D Q  
SEQ. ID. NO. 17 G Q K S E R K K W I H C F E N V I A L I Y L A S L S E Y D Q

SEQ. ID. NO. 16 C L E E N D Q E N R M E E S L A L F S T I L E L P W F K S T  
SEQ. ID. NO. 17 C L E E N N Q E N R M K E S L A L F G T I L E L P W F K S T

SEQ. ID. NO. 16 S V I L F L N K T D I L E D K I H T S H L A T Y F P S F Q G  
SEQ. ID. NO. 17 S V I L F L N K T D I L E E K I P T S H L A T Y F P S F Q G

SEQ. ID. NO. 16 P R R D A E A A K S F I L D M Y A R V Y A S C A E P Q D G G  
SEQ. ID. NO. 17 P K Q D A E A A K R F I L D M Y T R M Y T G C V D G P E G S

SEQ. ID. NO. 16 R K G S R A R R F F A H F T C A T D T Q S V R S V F K D V R  
SEQ. ID. NO. 17 K K G A R S R R L F S H Y T C A T D T Q N I R K V F K D V R

FIGURE 4b

SEQ. ID. NO. 16 D S V L A R Y L D E I N L L  
SEQ. ID. NO. 17 D S V L A R Y L D E I N L L

## ClustalW Formatted Alignments

SEQ. ID. NO. 18 A T G G C A T T T T A T A G C T G C T G C T G G G  
 SEQ. ID. NO. 19 A T G T T G C T G C T G C T A C T G G C G C  
 SEQ. ID. NO. 20 A T G G G G C C C G G G G C C C T T T T G C C C  
 SEQ. ID. NO. 21 A T G G C T T C C C C G C G G A G C T C C G G G C

SEQ. ID. NO. 18 T C C T C T T G G C A C T C A C C T G G C A C A C  
 SEQ. ID. NO. 19 C A C T C T T C C T C C G C C C C C G G G C G C  
 SEQ. ID. NO. 20 G G G T G G G G T G G C C A C T G C C G C T T C T  
 SEQ. ID. NO. 21 A G C C C G G G C C G C - G C C G C C G C C G C C

SEQ. ID. NO. 18 C T C T G C C T A C G G G C C A G A C C A G C G A  
 SEQ. ID. NO. 19 G G G C G G G G C G C A G A C C C C C A A C G C C  
 SEQ. ID. NO. 20 G G T T G T G A T G G C G G C A G G G G T G G C T  
 SEQ. ID. NO. 21 A C C G C C G C C C G C G C G C C T G C T A C T G

SEQ. ID. NO. 18 G C C C A A A A G A A G G G G G A C A T T A T C C  
 SEQ. ID. NO. 19 A C C T C A G A A G G T T G C C A G A T C A T A C  
 SEQ. ID. NO. 20 C C G G T G T G G G C C T C C C A C T C C C C C C  
 SEQ. ID. NO. 21 C T A C T G C T G C T G C C G C T G C T G C T G C

SEQ. ID. NO. 18 T T G G G G G G C T C T T T C C T A T T C A T T T  
 SEQ. ID. NO. 19 A C C C G C C C T G G G A A G G G G G C A T C A G  
 SEQ. ID. NO. 20 A T C T C C C G C G G C C T C A C T C G C G G G T  
 SEQ. ID. NO. 21 C T C T G G C G C C C G G G G C C T G G G G C T G

SEQ. ID. NO. 18 T G G A G T A G C A G C T A A A G A T C A A G A T  
 SEQ. ID. NO. 19 G T A C C G G G G C C T G A C T C G G G A C C A G  
 SEQ. ID. NO. 20 C C C C C G C A C C C C T C C T C A G A A C G G  
 SEQ. ID. NO. 21 G G C G C G G G G C G C C C C C C G G C C G C C G

SEQ. ID. NO. 18 C T C A A A T C A A G G C C G G A G T C T G T G G  
 SEQ. ID. NO. 19 G T G A A G G C T A T C A A C T T C C T G C C A G  
 SEQ. ID. NO. 20 C G C G C A G T G T A C A T C G G G G C A C T G T  
 SEQ. ID. NO. 21 C C C A G C A G C C C G C C G C T C T C C A T C A

SEQ. ID. NO. 18 A A T G T A T C A G G T A T A A T T T C C G T G G  
 SEQ. ID. NO. 19 T G G A C T A T G A G A T T G A G T A T G T G T G  
 SEQ. ID. NO. 20 T T C C C A T G A G C G G G G G C T G G C C A G G  
 SEQ. ID. NO. 21 T G G G C C T C A T G C C G C T C A C C A A G G A

Figure 5a

SEQ. ID. NO. 18 G T T T C G C T G G T T A C A G G C T A T G A T A  
 SEQ. ID. NO. 19 C C G G G G G G A G C G C G A G G T G G T G G G G  
 SEQ. ID. NO. 20 G G G C C A G G C C T G C C A G C C C G C G G T G  
 SEQ. ID. NO. 21 G G T G G C C A A G G G C A G C A T C G G G C G C

SEQ. ID. NO. 18 T T T G C C A T A G A G G A G A T A A A C A G C A  
 SEQ. ID. NO. 19 C C C A A G G T C C G C A A G T G C C T G G C C A  
 SEQ. ID. NO. 20 G A G A T G G C G C T G G A G G A C G T G A A T A  
 SEQ. ID. NO. 21 G G T G T G C T C C C C G C C G T G G A A C T G G

SEQ. ID. NO. 18 G C C C A G C C C T T C T T C C C A A C T T G A C  
 SEQ. ID. NO. 19 A C G G C T C C T G G A C A G A T A T G G A C A C  
 SEQ. ID. NO. 20 G C C G C A G G G A C A T C C T G C C G G A C T A  
 SEQ. ID. NO. 21 C C A T C G A G C A G A T C C G C A A C G A G T C

SEQ. ID. NO. 18 G C T G G G A T A C A G G A T A T T T G A C A C T  
 SEQ. ID. NO. 19 A C C C A G C C G C T G T G T C C G A A T C T G C  
 SEQ. ID. NO. 20 T G A G C T C A A G C T C A T C C A C C A C G A C  
 SEQ. ID. NO. 21 A C T C C T G C G C C C C T A C T T C C T C G A C

SEQ. ID. NO. 18 T G C A A C A C C G T T T C T A A G G C C T T G G  
 SEQ. ID. NO. 19 T C C A A G T C T T A T T T G A C C C T G G A A A  
 SEQ. ID. NO. 20 A G C A A G T G T G A T C C A G G C C A A G C C A  
 SEQ. ID. NO. 21 C T G C G G C T C T A T G A C A C G G A G T G C G

SEQ. ID. NO. 18 A A G C C A C C C T G A G T T T T G T T G C T C A  
 SEQ. ID. NO. 19 A T G G G A A G G T T T T C C T G A C G G G T G G  
 SEQ. ID. NO. 20 C C A A G T A C C T A T A T G A G C T G C T C T A  
 SEQ. ID. NO. 21 A C A A C G C A A A A G G G T T G A A A G C C T T

SEQ. ID. NO. 18 A A A C A A A A T T G A T T C T T T G A A C C T T  
 SEQ. ID. NO. 19 G G A C C T C C C A G C T C T G G A C G G A G C C  
 SEQ. ID. NO. 20 C A A C G A C C C T A T C A A G A T C A T C C T T  
 SEQ. ID. NO. 21 C T A C G A T G C A A T A A A A T A C G G G C C G

SEQ. ID. NO. 18 G A T G A G T T C T G C A A C T G C T C A G A G C  
 SEQ. ID. NO. 19 C G G G T G G A T T T C C G G T G T G A C C C C G  
 SEQ. ID. NO. 20 A T G C C T G G C T G C A G C T C T G T C T C C A  
 SEQ. ID. NO. 21 A A C C A C T T G A T G G T G T T T G G A G G C G

Figure 5b

SEQ. ID. NO. 18 A C A T T C C C T C T A C G A T T G C T G T G G T  
 SEQ. ID. NO. 19 A C T T C C A T C T G G T G G G C A G C T C C C G  
 SEQ. ID. NO. 20 C G C T G G T G G C T G A G G C T G C T A G G A T  
 SEQ. ID. NO. 21 T C T G T C C A T C C G T C A C A T C C A T C A T

SEQ. ID. NO. 18 G G G A G C A A C T G G C T C A G G C G T C T C C  
 SEQ. ID. NO. 19 G A G C A T C T G T A G T C A G G G C C A G T G G  
 SEQ. ID. NO. 20 G T G G A A C C T C A T T G T G C T T T C C T A T  
 SEQ. ID. NO. 21 T G C A G A G T C C C T C C A A G G C T G G A A T

SEQ. ID. NO. 18 A C G G C A G T G G C A A A T C T G C T G G G G C  
 SEQ. ID. NO. 19 A G C A C C C C C A A G C C C C A C T G C C A G G  
 SEQ. ID. NO. 20 G G C T C C A G C T C A C C A G C C C T G T C A A  
 SEQ. ID. NO. 21 C T G G T G C A G C T T T C T T T T G C T G C A A

SEQ. ID. NO. 18 T C T T C T A C A T T C C C C A G G T C A G T T A  
 SEQ. ID. NO. 19 T G A A T C G A A C G C C A C A C T C A G A A C G  
 SEQ. ID. NO. 20 A C C G G C A G C G T T T C C C C A C T T T C T T  
 SEQ. ID. NO. 21 C C A C G C C T G T T C T A G C C G A T A A G A A

SEQ. ID. NO. 18 T G C C T C C T C C A G C A G A C T C C T C A G C  
 SEQ. ID. NO. 19 G C G C G C A G T G T A C A T C G G G G C A C T G  
 SEQ. ID. NO. 20 C C G A A C G C A C C C A T C A G C C A C A C T C  
 SEQ. ID. NO. 21 A A A A T A C C C T T A T T T C T T T C G G A C C

SEQ. ID. NO. 18 A A C A A G A A T C A A T T C A A G T C T T T C C  
 SEQ. ID. NO. 19 T T T C C C A T G A G C G G G G G C T G G C C A G  
 SEQ. ID. NO. 20 C A C A A C C C T A C C C G C G T G A A A C T C T  
 SEQ. ID. NO. 21 G T C C C A T C A G A C A A T G C G G T G A A T C

SEQ. ID. NO. 18 T C C G A A C C A T C C C C A A T G A T G A G C A  
 SEQ. ID. NO. 19 G G G G C C A G G C C T G C C A G C C C G C G G T  
 SEQ. ID. NO. 20 T T G A A A A G T G G G G C T G G A A G A A G A T  
 SEQ. ID. NO. 21 C A G C C A T T C T G A A G T T G C T C A A G C A

SEQ. ID. NO. 18 C C A G G C C A C T G C C A T G G C A G A C A T C  
 SEQ. ID. NO. 19 G G A G A T G G C G C T G G A G G A C G T G A A T  
 SEQ. ID. NO. 20 T G C T A C C A T C C A G C A G A C C A C T G A G  
 SEQ. ID. NO. 21 C T A C C A G T G G A A G C G C G T G G G C A C G

Figure 5c

SEQ. ID. NO. 18 A T C G A G T A T T T C C G C T G G A A C T G G G  
 SEQ. ID. NO. 19 A G C C G C A G G G A C A T C C T G C C G G A C T  
 SEQ. ID. NO. 20 G T C T T C A C T T C G A C T C T G G A C G A C C  
 SEQ. ID. NO. 21 C T G A C G C A A G A C G T T C A G A G G T T C T

SEQ. ID. NO. 18 T G G G C A C A A T T G C A G C T G A T G A C G A  
 SEQ. ID. NO. 19 A T G A G C T C A A G C T C A T C C A C C A C G A  
 SEQ. ID. NO. 20 T G G A G G A A C G A G T G A A G G A G G C T G G  
 SEQ. ID. NO. 21 C T G A G G T G C G G A A T G A C C T G A C T G G

SEQ. ID. NO. 18 C T A T G G G C G G C C G G G G A T T G A G A A A  
 SEQ. ID. NO. 19 C A G C A A G T G T G A T C C A G G C C A A G C C  
 SEQ. ID. NO. 20 A A T T G A G A T T A C T T T C C G C C A G A G T  
 SEQ. ID. NO. 21 A G T T C T G T A T G G C G A G G A C A T T G A G

SEQ. ID. NO. 18 T T C C G A G A G G A A G C T G A G G A A A G G G  
 SEQ. ID. NO. 19 A C C A A G T A C C T A T A T G A G C T G C T C T  
 SEQ. ID. NO. 20 T T C T T C T C A G A T C C A G C T G T G C C C G  
 SEQ. ID. NO. 21 A T T T C A G A C A C C G A G A G C T T C T C C A

SEQ. ID. NO. 18 A T A T C T G C A T C G A C T T C A G T G A A C T  
 SEQ. ID. NO. 19 A C A A C G A C C C T A T C A A G A T C A T C C T  
 SEQ. ID. NO. 20 T C A A A A A C C T G A A G C G C C A G G A T G C  
 SEQ. ID. NO. 21 A C G A T C C C T G T A C C A G T G T C A A A A A

SEQ. ID. NO. 18 C A T C T C C C A G T A C T C T G A T G A G G A A  
 SEQ. ID. NO. 19 T A T G C C T G G C T G C A G C T C T G T C T C C  
 SEQ. ID. NO. 20 C C G A A T C A T C G T G G G A C T T T T C T A T  
 SEQ. ID. NO. 21 G C T G A A G G G G A A T G A T G T G C G G A T C

SEQ. ID. NO. 18 G A G A T C C A G C A T G T G G T A G A G G T G A  
 SEQ. ID. NO. 19 A C G C T G G T G G C T G A G G C T G C T A G G A  
 SEQ. ID. NO. 20 G A G A C T G A A G C C C G G A A A G T T T T T  
 SEQ. ID. NO. 21 A T C C T T G G C C A G T T T G A C C A G A A T A

SEQ. ID. NO. 18 T T C A A A A T T C C A C G G C C A A A G T C A T  
 SEQ. ID. NO. 19 T G T G G A A C C T C A T T G T G C T T T C C T A  
 SEQ. ID. NO. 20 G T G A G G T G T A C A A G G A G C G T C T C T T  
 SEQ. ID. NO. 21 T G G C A G C A A A A G T G T T C T G T T G T G C

Figure 5d

SEQ. ID. NO. 18 C G T G G T T T T C T C C A G T G G C C C A G A T  
 SEQ. ID. NO. 19 T G G C T C C A G C T C A C C A G C C C T G T C A  
 SEQ. ID. NO. 20 T G G G A A G A A G T A C G T C T G G T T C C T C  
 SEQ. ID. NO. 21 A T A C G A G G A G A A C A T G T A T G G T A G T

SEQ. ID. NO. 18 C T T G A G C C C C T C A T C A A G G A G A T T G  
 SEQ. ID. NO. 19 A A C C G G C A G C G T T T C C C C A C T T T C T  
 SEQ. ID. NO. 20 A T T G G G T G G T A T G C T G A C A A T T G G T  
 SEQ. ID. NO. 21 A A A T A T C A G T G G A T C A T T C C G G G C T

SEQ. ID. NO. 18 T C C G G C G C A A T A T C A C G G G C A A G A T  
 SEQ. ID. NO. 19 T C C G A A C G C A C C C A T C A G C C A C A C T  
 SEQ. ID. NO. 20 T C A A G A T C T A C G A C C C T T C T A T C A A  
 SEQ. ID. NO. 21 G G T A C G A G C C T T C T T G G T G G G A G C A

SEQ. ID. NO. 18 C T G G C T G G C C A G C G A G G C C T G G G C C  
 SEQ. ID. NO. 19 C C A C A A C C C T A C C C G C G T G A A A C T C  
 SEQ. ID. NO. 20 C T G C A C A G T G G A T G A G A T G A C T G A G  
 SEQ. ID. NO. 21 G G T G C A C A C G G A A G C C A A C T C A T C C

SEQ. ID. NO. 18 A G C T C C T C C C T G A T C G C C A T G C C T C  
 SEQ. ID. NO. 19 T T T G A A A A G T G G G G C T G G A A G A A G A  
 SEQ. ID. NO. 20 G C G G T G G A G G G C C A C A T C A C A A C T G  
 SEQ. ID. NO. 21 C G C T G C C T C C G G A A G A A T C T G C T T G

SEQ. ID. NO. 18 A G T A C T T C C A C G T G G T T G G C G G C A C  
 SEQ. ID. NO. 19 T T G C T A C C A T C C A G C A G A C C A C T G A  
 SEQ. ID. NO. 20 A G A T T G T C A T G C T G A A T C C T G C C A A  
 SEQ. ID. NO. 21 C T G C C A T G G A G G G C T A C A T T G G C G T

SEQ. ID. NO. 18 C A T T G G A T T C G C T C T G A A G G C T G G G  
 SEQ. ID. NO. 19 G G T C T T C A C T T C G A C T C T G G A C G A C  
 SEQ. ID. NO. 20 T A C C C G C A G C A T T T C C A A C A T G A C A  
 SEQ. ID. NO. 21 G G A T T T C G A G C C C C T G A G C T C C A A G

SEQ. ID. NO. 18 C A G A T C C C A G G C T T C C G G G A A T T C C  
 SEQ. ID. NO. 19 C T G G A G G A A C G A G T G A A G G A G G C T G  
 SEQ. ID. NO. 20 T C C C A G G A A T T T G T G G A G A A A C T A A  
 SEQ. ID. NO. 21 C A G A T C A A G A C C A T C T C A G G A A A G A

Figure 5e



SEQ. ID. NO. 18 T G A A G A A G G T C C A T C C C A G G A A G T C  
 SEQ. ID. NO. 19 G A A T T G A G A T T A C T T T C C G C C A G A G  
 SEQ. ID. NO. 20 C C A A G C G A C T G A A A A G A C A C C C T G A  
 SEQ. ID. NO. 21 C T C C A C A G C A G T A T G A G A G A G A G T A

SEQ. ID. NO. 18 T G T C C A C A A T G G T T T T G C C A A G G A G  
 SEQ. ID. NO. 19 T T T C T T C T C A G A T C C A G C T G T G C C C  
 SEQ. ID. NO. 20 G G A G A C A G G A G G C T T C C A G G A G G C A  
 SEQ. ID. NO. 21 C A A C A A C A A G C G G T C A G G C G T G G G G

SEQ. ID. NO. 18 T T T T G G G A A G A A A C A T T T A A C T G C C  
 SEQ. ID. NO. 19 G T C A A A A A C C T G A A G C G C C A G G A T G  
 SEQ. ID. NO. 20 C C G C T G G C C T A T G A T G C C A T C T G G G  
 SEQ. ID. NO. 21 C C C A G C A A G T T C C A C G G G T A C G C C T

SEQ. ID. NO. 18 A C C T C C A A G A A G G T G C A A A A G G A C C  
 SEQ. ID. NO. 19 C C C G A A T C A T C G T G G G A C T T T T C T A  
 SEQ. ID. NO. 20 C C T T G G C A C T G G C C C T G A A C A A G A C  
 SEQ. ID. NO. 21 A C G A T G G C A T C T G G G T C A T C G C C A A

SEQ. ID. NO. 18 T T T A C C T G T G G A C A C C T T T C T G A G A  
 SEQ. ID. NO. 19 T G A G A C T G A A G C C C G G A A A G T T T T T  
 SEQ. ID. NO. 20 A T C T G G A G G A G G C G G C C G T T C T G G T  
 SEQ. ID. NO. 21 G A C A C T G C A G A G G G C C A T G G A G A C A

SEQ. ID. NO. 18 G G T C A C G A A G A A A G T G G C G A C A G G T  
 SEQ. ID. NO. 19 T G T G A G G T G T A C A A G G A G C G T C T C T  
 SEQ. ID. NO. 20 G T G C G C C T G G A G G A C T T C A A C T A C A  
 SEQ. ID. NO. 21 C T G C A T G C C A G C A G C C G G C A C C A G C

SEQ. ID. NO. 18 T T A G C A A C A G C T C G A C A G C C T T C C G  
 SEQ. ID. NO. 19 T T G G G A A G A A G T A C G T C T G G T T C C T  
 SEQ. ID. NO. 20 A C A A C C A G A C C A T T A C C G A C C A A A T  
 SEQ. ID. NO. 21 G G A T C C A G G A C T T C A A C T A C A C G G A

SEQ. ID. NO. 18 A C C C C T C T G T A C A G G G G A T G A G A A C  
 SEQ. ID. NO. 19 C A T T G G G T G G T A T G C T G A C A A T T G G  
 SEQ. ID. NO. 20 C T A C C G G G C A A T G A A C T C T T C G T C C  
 SEQ. ID. NO. 21 C C A C A C G C T G G G C A G G A T C A T C C T C

Figure 5f

SEQ. ID. NO. 18 A T C A G C A G T G T C G A G A C C C C T T A C A  
 SEQ. ID. NO. 19 T T C A A G A T C T A C G A C C C T T C T A T C A  
 SEQ. ID. NO. 20 T T T G A G G G T G T C T C T G G C C A T G T G G  
 SEQ. ID. NO. 21 A A T G C C A T G A A C G A G A C C A A C T T C T

SEQ. ID. NO. 18 T A G A T T A C A C G C A T T T A C G G A T A T C  
 SEQ. ID. NO. 19 A C T G C A C A G T G G A T G A G A T G A C T G A  
 SEQ. ID. NO. 20 T G T T T G A T G C C A G C G G C T C T C G G A T  
 SEQ. ID. NO. 21 T C G G G G T C A C G G G T C A A G T T G T A T T

SEQ. ID. NO. 18 C T A C A A T G T G T A C T T A G C A G T C T A C  
 SEQ. ID. NO. 19 G G C G G T G G A G G G C C A C A T C A C A A C T  
 SEQ. ID. NO. 20 G G C A T G G A C G C T T A T C G A G C A G C T T  
 SEQ. ID. NO. 21 C C G G A A T G G G G A G A G A A T G G G G A C C

SEQ. ID. NO. 18 T C C A T T G C C C A C G C C T T G C A A G A T A  
 SEQ. ID. NO. 19 G A G A T T G T C A T G C T G A A T C C T G C C A  
 SEQ. ID. NO. 20 C A G G G T G G C A G C T A C A A G A A G A T T G  
 SEQ. ID. NO. 21 A T T A A A T T T A C T C A A T T T C A A G A C A

SEQ. ID. NO. 18 T A T A T A C C T G C T T A C C T G G G A G A G G  
 SEQ. ID. NO. 19 A T A C C C G C A G C A T T T T C C A A C A T G A C  
 SEQ. ID. NO. 20 G C T A C T A T G A C A G C A C C A A G G A T G A  
 SEQ. ID. NO. 21 G C A G G G A G G T G A A G G T G G G A G A G T A

SEQ. ID. NO. 18 G C T C T T C A C C A A T G G C T C C T G T G C A  
 SEQ. ID. NO. 19 A T C C C A G G A A T T T G T G G A G A A A C T A  
 SEQ. ID. NO. 20 T C T T T C C T G G T C C A A A A C A G A T A A A  
 SEQ. ID. NO. 21 C A A C G C T G T G G C C G A C A C A C T G G A G

SEQ. ID. NO. 18 G A C A T C A A G A A A G T T G A G G C G T G G C  
 SEQ. ID. NO. 19 A C C A A G C G A C T G A A A A G A C A C C C T G  
 SEQ. ID. NO. 20 T G G A T T G G A G G G T C C C C C C C A G C T G  
 SEQ. ID. NO. 21 A T C A T C A A T G A C A C C A T C A G G T T C C

SEQ. ID. NO. 18 A G G T C C T G A A G C A C C T A C G G C A T C T  
 SEQ. ID. NO. 19 A G G A G A C A G G A G G C T T C C A G G A G G C  
 SEQ. ID. NO. 20 A C C A G A C C C T G G T C A T C A A G A C A T T  
 SEQ. ID. NO. 21 A A G G A T C C G A A C C A C C A A A A G A C A A

Figure 5g

SEQ. ID. NO. 18 A A A C T T T A C A A A C A A T A T G G G G G A G  
 SEQ. ID. NO. 19 A C C G C T G G C C T A T G A T G C C A T C T G G  
 SEQ. ID. NO. 20 C C G C T T C C T G T C A C A G A A A C T C T T T  
 SEQ. ID. NO. 21 G A C C A T C A T C C T G G A G C A G C T G C G G

SEQ. ID. NO. 18 C A G G T G A C C T T T G A T G A G T G T G G T G  
 SEQ. ID. NO. 19 G C C T T G G C A C T G G C C C T G A A C A A G A  
 SEQ. ID. NO. 20 A T C T C C G T C T C A G T T C T C T C C A G C C  
 SEQ. ID. NO. 21 A A G A T C T C C C T A C C T C T C T A C A G C A

SEQ. ID. NO. 18 A C C T G G T G G G G A A C T A T T C C A T C A T  
 SEQ. ID. NO. 19 C A T C T G G A G G A G G C G G C C G T T C T G G  
 SEQ. ID. NO. 20 T G G G C A T T G T C C T A G C T G T T G T C T G  
 SEQ. ID. NO. 21 T C C T C T C T G C C C T C A C C A T C C T C G G

SEQ. ID. NO. 18 C A A C T G G C A C C T C T C C C C A G A G G A T  
 SEQ. ID. NO. 19 T G T G C G C C T G G A G G A C T T C A A C T A C  
 SEQ. ID. NO. 20 T C T G T C C T T T A A C A T C T A C A A C T C A  
 SEQ. ID. NO. 21 G A T G A T C A T G G C C A G T G C T T T T C T C

SEQ. ID. NO. 18 G G C T C C A T C G T G T T T A A G G A A G T C G  
 SEQ. ID. NO. 19 A A C A A C C A G A C C A T T A C C G A C C A A A  
 SEQ. ID. NO. 20 C A T G T C C G T T A T A T C C A G A A C T C A C  
 SEQ. ID. NO. 21 T T C T T C A A C A T C A A G A A C C G G A A T C

SEQ. ID. NO. 18 G G T A T T A C A A C G T C T A T G C C A A G A A  
 SEQ. ID. NO. 19 T C T A C C G G G C A A T G A A C T C T T C G T C  
 SEQ. ID. NO. 20 A G C C C A A C C T G A A C A A C C T G A C T G C  
 SEQ. ID. NO. 21 A G A A G C T C A T A A A G A T G T C G A G T C C

SEQ. ID. NO. 18 G G G A G A A A G A C T C T T C A T C A A C G A G  
 SEQ. ID. NO. 19 C T T T G A G G G T G T C T C T G G C C A T G T G  
 SEQ. ID. NO. 20 T G T G G G C T G C T C A C T G G C T T T A G C T  
 SEQ. ID. NO. 21 A T A C A T G A A C A A C C T T A T C A T C C T T

SEQ. ID. NO. 18 G A G A A A A T C C T G T G G A G T G G G T T C T  
 SEQ. ID. NO. 19 G T G T T T G A T G C C A G C G G C T C T C G G A  
 SEQ. ID. NO. 20 G C T G T C T T C C C C C T G G G G C T C G A T G  
 SEQ. ID. NO. 21 G G A G G G A T G C T C T C C T A T G C T T C C A

Figure 5h

SEQ. ID. NO. 18 C C A G G G A G G T G C C C T T C T C C A A C T G  
 SEQ. ID. NO. 19 T G G C A T G G A C G C T T A T C G A G C A G C T  
 SEQ. ID. NO. 20 G T T A C C A C A T T G G G A G G A A C C A G T T  
 SEQ. ID. NO. 21 T A T T T C T C T T T G G C C T T G A T G G A T C

SEQ. ID. NO. 18 C A G C C G A G A C T G C C T G G C A G G G A C C  
 SEQ. ID. NO. 19 T C A G G G T G G C A G C T A C A A G A A G A T T  
 SEQ. ID. NO. 20 T C C T T T C G T C T G C C A G G C C C G C C T C  
 SEQ. ID. NO. 21 C T T T G T C T C T G A A A A G A C C T T T G A A

SEQ. ID. NO. 18 A G G A A A G G G A T C A T T G A G G G G G A G C  
 SEQ. ID. NO. 19 G G C T A C T A T G A C A G C A C C A A G G A T G  
 SEQ. ID. NO. 20 T G G C T C C T G G G C C T G G G C T T T A G T C  
 SEQ. ID. NO. 21 A C A C T T T G C A C C G T C A G G A C C T G G A

SEQ. ID. NO. 18 C C A C C T G C T G C T T T G A G T G T G T G G A  
 SEQ. ID. NO. 19 A T C T T T C C T G G T C C A A A A C A G A T A A  
 SEQ. ID. NO. 20 T G G G C T A C G G T T C C A T G T T C A C C A A  
 SEQ. ID. NO. 21 T T C T C A C C G T G G G C T A C A C G A C C G C

SEQ. ID. NO. 18 G T G T C C T G A T G G G G A G T A T A G T G A T  
 SEQ. ID. NO. 19 A T G G A T T G G A G G G T C C C C C C C A G C T  
 SEQ. ID. NO. 20 G A T T T G G T G G G T C C A C A C G G T C T T C  
 SEQ. ID. NO. 21 T T T T G G G G C C A T G T T T G C A A A G A C C

SEQ. ID. NO. 18 G A G A C A G A T G C C A G T G C C T G T A A C A  
 SEQ. ID. NO. 19 G A C C A G A C C C T G G T C A T C A A G A C A T  
 SEQ. ID. NO. 20 A C A A A G A A G G A A G A A A A G A A G G A G T  
 SEQ. ID. NO. 21 T G G A G A G T C C A C G C C A T C T T C A A A A

SEQ. ID. NO. 18 A G T G C C C A G A T G A C T T C T G G T C C A A  
 SEQ. ID. NO. 19 T C C G C T T C C T G T C A C A G A A A C T C T T  
 SEQ. ID. NO. 20 G G A G G A A G A C T C T G G A A C C C T G G A A  
 SEQ. ID. NO. 21 A T G T G A A A A T G A A G A A G A A G A T C A T

SEQ. ID. NO. 18 T G A G A A C C A C A C C T C C T G C A T T G C C  
 SEQ. ID. NO. 19 T A T C T C C G T C T C A G T T C T C T C C A G C  
 SEQ. ID. NO. 20 G C T G T A T G C C A C A G T G G G C C T G C T G  
 SEQ. ID. NO. 21 C A A G G A C C A G A A A C T G C T T G T G A T C

Figure 5i

SEQ. ID. NO. 18 A A G G A G A T C G A G T T T C T G T C G T G G A  
 SEQ. ID. NO. 19 C T G G G C A T T G T C C T A G C T G T T G T C T  
 SEQ. ID. NO. 20 G T G G G C A T G G A T G T C C T C A C T C T C G  
 SEQ. ID. NO. 21 G T G G G G G G C A T G C T G C T G A T C G A C C

SEQ. ID. NO. 18 C G G A G C C C T T T G G G A T C G C A C T C A C  
 SEQ. ID. NO. 19 G T C T G T C C T T T A A C A T C T A C A A C T C  
 SEQ. ID. NO. 20 C C A T C T G G C A G A T C G T G G A C C C T C T  
 SEQ. ID. NO. 21 T G T G T A T C C T G A T C T G C T G G C A G G C

SEQ. ID. NO. 18 C C T C T T T G C C G T G C T G G G C A T T T T T C  
 SEQ. ID. NO. 19 A C A T G T C C G T T A T A T C C A G A A C T C A  
 SEQ. ID. NO. 20 G C A C C G G A C C A T T G A G A C A T T T G C C  
 SEQ. ID. NO. 21 T G T G G A C C C C C T G C G A A G G A C A G T G

SEQ. ID. NO. 18 C T G A C A G C C T T T G T G C T G G G T G T G T  
 SEQ. ID. NO. 19 C A G C C C A A C C T G A A C A A C C T G A C T G  
 SEQ. ID. NO. 20 A A G G A G G A A C C T A A G G A A G A T A T T G  
 SEQ. ID. NO. 21 G A G A A G T A C A G C A T G G A G C C G G A C C

SEQ. ID. NO. 18 T T A T C A A G T T C C G C A A C A C A C C C C A T  
 SEQ. ID. NO. 19 C T G T G G G C T G C T C A C T G G C T T T A G C  
 SEQ. ID. NO. 20 A C G T C T C T A T T C T G C C C C A G C T G G A  
 SEQ. ID. NO. 21 C A G C A G G A C G G G A T A T C T C C A T C C G

SEQ. ID. NO. 18 T G T C A A G G C C A C C A A C C G A G A G C T C  
 SEQ. ID. NO. 19 T G C T G T C T T C C C C C T G G G G C T C G A T  
 SEQ. ID. NO. 20 G C A T T G C A G C T C C A G G A A G A T G A A T  
 SEQ. ID. NO. 21 C C C T C T C C T G G A G C A C T G T G A G A A C

SEQ. ID. NO. 18 T C C T A C C T C C T C C T C T T C T C C C T G C  
 SEQ. ID. NO. 19 G G T T A C C A C A T T G G G A G G A A C C A G T  
 SEQ. ID. NO. 20 A C A T G G C T T G G C A T T T T C T A T G G T T  
 SEQ. ID. NO. 21 A C C C A T A T G A C C A T C T G G C T T G G C A

SEQ. ID. NO. 18 T C T G C T G C T T C T C C A G C T C C C T G T T  
 SEQ. ID. NO. 19 T T C C T T T C G T C T G C C A G G C C C G C C T  
 SEQ. ID. NO. 20 A C A A G G G G C T G C T G C T G C T G C T G G G  
 SEQ. ID. NO. 21 T C G T C T A T G C C T A C A A G G G A C T T C T

Figure 5J

SEQ. ID. NO. 18 C T T C A T C G G G G A G C C C C A G G A C T G G  
 SEQ. ID. NO. 19 C T G G C T C C T G G G C C T G G G C T T T A G T  
 SEQ. ID. NO. 20 A A T C T T C C T T G C T T A T G A G A C C A A G  
 SEQ. ID. NO. 21 C A T G T T G T T C G G T T G T T T C T T A G C T

SEQ. ID. NO. 18 A C G T G C C G C C T G C G C C A G C C G G C C T  
 SEQ. ID. NO. 19 C T G G G C T A C G G T T C C A T G T T C A C C A  
 SEQ. ID. NO. 20 A G T G T G T C C A C T G A G A A G A T C A A T G  
 SEQ. ID. NO. 21 T G G G A G A C C C G C A A C G T C A G C A T C C

SEQ. ID. NO. 18 T T G G C A T C A G C T T C G T G C T C T G C A T  
 SEQ. ID. NO. 19 A G A T T T G G T G G G T C C A C A C G G T C T T  
 SEQ. ID. NO. 20 A T C A C C G G G C T G T G G G C A T G G C T A T  
 SEQ. ID. NO. 21 C C G C A C T C A A C G A C A G C A A G T A C A T

SEQ. ID. NO. 18 C T C A T G C A T C C T G G T G A A A A C C A A C  
 SEQ. ID. NO. 19 C A C A A A G A A G G A A G A A A A G A A G G A G  
 SEQ. ID. NO. 20 C T A C A A T G T G G C A G T C C T G T G C C T C  
 SEQ. ID. NO. 21 C G G G A T G A G T G T C T A C A A C G T G G G G

SEQ. ID. NO. 18 C G T G T C C T C C T G G T G T T T G A G G C C A  
 SEQ. ID. NO. 19 T G G A G G A A G A C T C T G G A A C C C T G G A  
 SEQ. ID. NO. 20 A T C A C T G C T C C T G T C A C C A T G A T T C  
 SEQ. ID. NO. 21 A T C A T G T G C A T C A T C G G G G C C G C T G

SEQ. ID. NO. 18 A G A T C C C C A C C A G C T T C C A C C G C A A  
 SEQ. ID. NO. 19 A G C T G T A T G C C A C A G T G G G C C T G C T  
 SEQ. ID. NO. 20 T G T C C A G C C A G C A G G A T G C A G C C T T  
 SEQ. ID. NO. 21 T C T C C T T C C T G A C C C G G G A C C A G C C

SEQ. ID. NO. 18 G T G G T G G G G G C T C A A C C T G C A G T T C  
 SEQ. ID. NO. 19 G G T G G G C A T G G A T G T C C T C A C T C T C  
 SEQ. ID. NO. 20 T G C C T T T G C C T C T C T T G C C A T A G T T  
 SEQ. ID. NO. 21 C A A T G T G C A G T T C T G C A T C G T G G C T

SEQ. ID. NO. 18 C T G C T G G T T T T C C T C T G C A C C T T C A  
 SEQ. ID. NO. 19 G C C A T C T G G C A G A T C G T G G A C C C T C  
 SEQ. ID. NO. 20 T T C T C C T C C T A T A T C A C T C T T G T T G  
 SEQ. ID. NO. 21 C T G G T C A T C A T C T T C T G C A G C A C C A

Figure 5k

SEQ. ID. NO. 18 T G C A G A T T G T C A T C T G T G T G A T C T G  
 SEQ. ID. NO. 19 T G C A C C G G A C C A T T G A G A C A T T T G C  
 SEQ. ID. NO. 20 T G C T C T T T G T G C C C A A G A T G C G C A G  
 SEQ. ID. NO. 21 T C A C C C T C T G C C T G G T A T T C G T G C C

SEQ. ID. NO. 18 G C T C T A C A C C G C G C C C C C C T C A A G C  
 SEQ. ID. NO. 19 C A A G G A G G A A C C T A A G G A A G A T A T T  
 SEQ. ID. NO. 20 G C T G A T C A C C C G A G G G G A A T G G C A G  
 SEQ. ID. NO. 21 G A A G C T C A T C A C C C T G A G A A C A A A C

SEQ. ID. NO. 18 T A C C G C A A C C A G G A G C T G G A G G A T G  
 SEQ. ID. NO. 19 G A C G T C T C T A T T C T G C C C C A G C T G G  
 SEQ. ID. NO. 20 T C G G A G G C G C A G G A C A C C A T G A A G A  
 SEQ. ID. NO. 21 C C A G A T G C A G C A A C G C A G A A C A G G C

SEQ. ID. NO. 18 A G A T C A T C T T C A T C A C G T G C C A C G A  
 SEQ. ID. NO. 19 A G C A T T G C A G C T C C A G G A A G A T G A A  
 SEQ. ID. NO. 20 C A G G G T C A T C G A C C A A C A A C A C G A  
 SEQ. ID. NO. 21 G A T T C C A G T T C A C T C A G A A T C A G A A

SEQ. ID. NO. 18 G G G C T C C C T C A T G G C C C T G G G C T T C  
 SEQ. ID. NO. 19 T A C A T G G C T T G G C A T T T T C T A T G G T  
 SEQ. ID. NO. 20 G G A G G A G A A G T C C C G G C T G T T G G A G  
 SEQ. ID. NO. 21 G A A A G A A G A T T C T A A A A C G T C C A C C

SEQ. ID. NO. 18 C T G A T C G G C T A C A C C T G C C T G C T G G  
 SEQ. ID. NO. 19 T A C A A G G G G C T G C T G C T G C T G C T G G  
 SEQ. ID. NO. 20 A A G G A G A A C C G T G A A C T G G A A A A G A  
 SEQ. ID. NO. 21 T C G G T C A C C A G T G T G A A C C A A G C C A

SEQ. ID. NO. 18 C T G C C A T C T G C T T C T T C T T T G C C T T  
 SEQ. ID. NO. 19 G A A T C T T C C T T G C T T A T G A G A C C A A  
 SEQ. ID. NO. 20 T C A T T G C T G A G A A A G A G G A G C G T G T  
 SEQ. ID. NO. 21 G C A C A T C C C G C C T G G A G G G C C T A C A

SEQ. ID. NO. 18 C A A G T C C C G G A A G C T G C C G G A G A A C  
 SEQ. ID. NO. 19 G A G T G T G T C C A C T G A G A A G A T C A A T  
 SEQ. ID. NO. 20 C T C T G A A C T G C G C C A T C A A C T C C A G  
 SEQ. ID. NO. 21 G T C A G A A A A C C A T C G C C T G C G A A T G

Figure 51

SEQ. ID. NO. 18 T T C A A T G A A G C C A A G T T C A T C A C C T  
 SEQ. ID. NO. 19 G A T C A C C G G G C T G T G G G C A T G G C T A  
 SEQ. ID. NO. 20 T C T C G G C A G C A G C T C C G C T C C C G G C  
 SEQ. ID. NO. 21 A A G A T C A C A G A G C T G G A T A A A G A C T

SEQ. ID. NO. 18 T C A G C A T G C T C A T C T T C T T C A T C G T  
 SEQ. ID. NO. 19 T C T A C A A T G T G G C A G T C C T G T G C C T  
 SEQ. ID. NO. 20 G C C A C C C A C C G A C A C C C C C A G A A C C  
 SEQ. ID. NO. 21 T G G A A G A G G T C A C C A T G C A G C T G C A

SEQ. ID. NO. 18 C T G G A T C T C C T T C A T T C C A G C C T A T  
 SEQ. ID. NO. 19 C A T C A C T G C T C C T G T C A C C A T G A T T  
 SEQ. ID. NO. 20 C T C T G G G G G C C T G C C C A G G G G A C C C  
 SEQ. ID. NO. 21 G G A C A C A C C A G A A A A G A C C A C C T A C

SEQ. ID. NO. 18 G C C A G C A C C T A T G G C A A G T T T G T C T  
 SEQ. ID. NO. 19 C T G T C C A G C C A G C A G G A T G C A G C C T  
 SEQ. ID. NO. 20 C C T G A G C C C C C C G A C C G G C T T A G C T  
 SEQ. ID. NO. 21 A T T A A A C A G A A C C A C T A C C A A G A G C

SEQ. ID. NO. 18 C T G C C G T A G A G G T G A T T G C C A T C C T  
 SEQ. ID. NO. 19 T T G C C T T T G C C T C T C T T G C C A T A G T  
 SEQ. ID. NO. 20 G T G A T G G G A G T C G A G T G C A T T T G C T  
 SEQ. ID. NO. 21 T C A A T G A C A T C C T C A A C C T G G G A A A

SEQ. ID. NO. 18 G G C A G C C A G C T T T G G C T T G C T G G C G  
 SEQ. ID. NO. 19 T T T C T C C T C C T A T A T C A C T C T T G T T  
 SEQ. ID. NO. 20 T T A T A A G T G A G G G T A G G G T G A G G G A  
 SEQ. ID. NO. 21 C T T C A C T G A G A G C A C A G A T G G A G G A

SEQ. ID. NO. 18 T G C A T C T T C T T C A A C A A G A T C T A C A  
 SEQ. ID. NO. 19 G T G C T C T T T G T G C C C A A G A T G C G C A  
 SEQ. ID. NO. 20 G G A C A G G C C A G T A G G G G G A G G G A A A  
 SEQ. ID. NO. 21 A A G G C C A T T T T A A A A A A T C A C C T C G

SEQ. ID. NO. 18 T C A T T C T C T T C A A G C C A T C C C G C A A  
 SEQ. ID. NO. 19 G G C T G A T C A C C C G A G G G G A A T G G C A  
 SEQ. ID. NO. 20 G G G A G A G G G G A A G G G C A G G G A C T C  
 SEQ. ID. NO. 21 A T C A A A A T C C C C A G C T A C A G T G G A A

Figure 5m



SEQ. ID. NO. 18 C A C C A T C G A G G A G G T G C G T T G C A G C  
 SEQ. ID. NO. 19 G T C G G A G G C G C A G G A C A C C A T G A A G  
 SEQ. ID. NO. 20 A G G A A G C A G G G G G T C C C C A T C C C C A  
 SEQ. ID. NO. 21 C A C A A C A G A G C C C T C T C G A A C A T G C

SEQ. ID. NO. 18 A C C G C A G C T C A C G C T T T C A A G G T G G  
 SEQ. ID. NO. 19 A C A G G G T C A T C G A C C A A C A A C A A C G  
 SEQ. ID. NO. 20 G C T G G G A A G A A C A T G C T A T C C A A T C  
 SEQ. ID. NO. 21 A A A G A T C C T A T A G A A G A T A T A A A C T

SEQ. ID. NO. 18 C T G C C C G G G C C A C G C T G C G C C G C A G  
 SEQ. ID. NO. 19 A G G A G G A G A A G T C C C G G C T G T T G G A  
 SEQ. ID. NO. 20 T C A T C T C T T G T A A A T A C A T G T C C C C  
 SEQ. ID. NO. 21 C T C C A G A A C A C A T C C A G C G T C G G C T

SEQ. ID. NO. 18 C A A C G T C T C C C G C A A G C G G T C C A G C  
 SEQ. ID. NO. 19 G A A G G A G A A C C G T G A A C T G G A A A A G  
 SEQ. ID. NO. 20 C T G T G A G T T C T G G G C T G A T T T G G G T  
 SEQ. ID. NO. 21 G T C C C T C C A G C T C C C C A T C C T C C A C

SEQ. ID. NO. 18 A G C C T T G G A G G C T C C A C G G G A T C C A  
 SEQ. ID. NO. 19 A T C A T T G C T G A G A A A G A G G A G C G T G  
 SEQ. ID. NO. 20 C T C T C A T A C C T C T G G G A A A C A G A C C  
 SEQ. ID. NO. 21 C A C G C C T A C C T C C C A T C C A T C G G A G

SEQ. ID. NO. 18 C C C C C T C C T C C T C C A T C A G C A G C A A  
 SEQ. ID. NO. 19 T C T C T G A A C T G C G C C A T C A G C T C C A  
 SEQ. ID. NO. 20 T T T T T C T C T C T T A C T G C T T C A T G T A  
 SEQ. ID. NO. 21 G C G T G G A C G C C A G C T G T G T C A G C C C

SEQ. ID. NO. 18 G A G C A A C A G C G A A G A C C C A T T C C C A  
 SEQ. ID. NO. 19 G T C T C G G C A G C A G C T C C G C T C C C G G  
 SEQ. ID. NO. 20 A T T T T G T A T C A C C T C T T C A C A A T T T  
 SEQ. ID. NO. 21 C T G C G T C A G C C C C A C C G C C A G C C C C

SEQ. ID. NO. 18 C A G C C C G A G A G G C A G A A G C A G C A G C  
 SEQ. ID. NO. 19 C G C C A C C C A C C G A C A C C C C C A G A A C  
 SEQ. ID. NO. 20 A G T T C G T A C C T G G C T T G A A G C T G C T  
 SEQ. ID. NO. 21 C G C C A C A G A C A T G T G C C A C C C T C C T

Figure 5n

SEQ. ID. NO. 18 A G C C G C T G G C C C T A A C C C A G C A A G A  
 SEQ. ID. NO. 19 C C T C T G G G G G C C T G C C C A G G G G A C C  
 SEQ. ID. NO. 20 C A C T G C T C A C A C G C T G C C T C C T C A G  
 SEQ. ID. NO. 21 T C C G A G T C A T G G T C T C G G G C C T G T A

SEQ. ID. NO. 18 G C A G C A G C A G C A G C C C C T G A C C C T C  
 SEQ. ID. NO. 19 C C C T G A G C C C C C G A C C G G C T T A G C  
 SEQ. ID. NO. 20 C A G C C T C A C T G C A T C T T T C T C T T C C  
 SEQ. ID. NO. 21 A G G G T G G G A G G C C T G G G C C C G G G G C

SEQ. ID. NO. 18 C C A C A G C A G C A A C G A T C T C A G C A G C  
 SEQ. ID. NO. 19 T G T G A T G G G A G T C G A G T G C A T T T G C  
 SEQ. ID. NO. 20 C A T G C A A C A C C C T C T T C T A G T T A C C  
 SEQ. ID. NO. 21 C T C C C C C G T G A C A G A A C C A C A C T G G

SEQ. ID. NO. 18 A G C C C A G A T G C A A G C A G A A G G T C A T  
 SEQ. ID. NO. 19 T T T A T A A G T G A G G G T A G G G T G A G G G  
 SEQ. ID. NO. 20 A C G G C A A C C C C T  
 SEQ. ID. NO. 21 G C A G A G G G G T C T G C T G C A G A A A C A C

SEQ. ID. NO. 18 C T T T G G C A G C G G C A C G G T C A C C T T C  
 SEQ. ID. NO. 19 A G G A C A G G C C A G T A G G G G G A G G G A A  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21 T G T C G G C T C T G G C T G C G G A G A A G C T

SEQ. ID. NO. 18 T C A C T G A G C T T T G A T G A G C C T C A G A  
 SEQ. ID. NO. 19 A G G G A G A G G G G A A G G G C A G G G G A C T  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21 G G G C A C C A T G G C T G G C C T C T C A G G A

SEQ. ID. NO. 18 A G A A C G C C A T G G C C C A C G G G A A T T C  
 SEQ. ID. NO. 19 C A G G A A G C A G G G G G T C C C C A T C C C C  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21 C C A C T C G G A T G G C A C T C A G G T G G A C

SEQ. ID. NO. 18 T A C G C A C C A G A A C T C C C T G G A G G C C  
 SEQ. ID. NO. 19 A G C T G G G A A G A A C A T G C T A T C C A A T  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21 A G G A C G G G G C A G G G G G A G A C T T G G C

Figure 5o

SEQ. ID. NO. 18 C A G A A A A G C A G C G A T A C G C T G A C C C  
 SEQ. ID. NO. 19 C T C A T C T C T T G T A A A T A C A T G T C C C  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21 A C C T G A C C T C G A G C C T T A T T T G T G A

SEQ. ID. NO. 18 G A C A C C A G C C A T T A C T C C C G C T G C A  
 SEQ. ID. NO. 19 C C T G T G A G T T C T G G G C T G A T T T G G G  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21 A G T C C T T A T T T C T T C A C A A G A A G A

SEQ. ID. NO. 18 G T G C G G G G A A A C G G A C T T A G A T C T G  
 SEQ. ID. NO. 19 T C T C T C A T A C C T C T G G G A A A C A G A C  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21 G G A A C G G A A A T G G G A C G T C T T C C T T

SEQ. ID. NO. 18 A C C G T C C A G G A A A C A G G T C T G C A A G  
 SEQ. ID. NO. 19 C T T T T T C T C T C T T A C T G C T T C A T G T  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21 A A C A T C T G C A A A C A A G G A G G C G C T G

SEQ. ID. NO. 18 G A C C T G T G G G T G G A G A C C A G C G G C C  
 SEQ. ID. NO. 19 A A T T T T G T A T C A C C T C T T C A C A A T T  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21 G G A T A T C A A A C T T G C A A A A A A A A A A

SEQ. ID. NO. 18 A G A G G T G G A G G A C C C T G A A G A G T T G  
 SEQ. ID. NO. 19 T A G T T C G T A C C T G G C T T G A A G C T G C  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21 A A A A A A A A A A A A

SEQ. ID. NO. 18 T C C C C A G C A C T T G T A G T G T C C A G T T  
 SEQ. ID. NO. 19 T C A C T G C T C A C A C G C T G C C T C C T C A  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21

SEQ. ID. NO. 18 C A C A G A G C T T T G T C A T C A G T G G T G G  
 SEQ. ID. NO. 19 G C A G C C T C A C T G C A T C T T T C T C T T C  
 SEQ. ID. NO. 20  
 SEQ. ID. NO. 21

Figure 5p

SEQ. ID. NO. 18 A G G C A G C A C T G T T A C A G A A A C G T A  
SEQ. ID. NO. 19 C C A T G C A A C A C C C T C T T C T A G T T A C  
SEQ. ID. NO. 20  
SEQ. ID. NO. 21

SEQ. ID. NO. 18 G T G A A T T C A  
SEQ. ID. NO. 19 C A C G G C A A C C C C T G C A G C T C C T C T G  
SEQ. ID. NO. 20  
SEQ. ID. NO. 21

SEQ. ID. NO. 18  
SEQ. ID. NO. 19 C C T T T G T G C T C T G T T C C T G T C C A G C  
SEQ. ID. NO. 20  
SEQ. ID. NO. 21

SEQ. ID. NO. 18  
SEQ. ID. NO. 19 A G G G G T C T C C C A A C A A G T G C T C T T T  
SEQ. ID. NO. 20  
SEQ. ID. NO. 21

SEQ. ID. NO. 18  
SEQ. ID. NO. 19 C C A C C C C A A A G G G G C C T C T C C T T T T  
SEQ. ID. NO. 20  
SEQ. ID. NO. 21

SEQ. ID. NO. 18  
SEQ. ID. NO. 19 C T C C A C T G T C A T A A T C T C T T T C C A T  
SEQ. ID. NO. 20  
SEQ. ID. NO. 21

SEQ. ID. NO. 18  
SEQ. ID. NO. 19 C T T A C T T G C C C T T C T A T A C T T T C T C  
SEQ. ID. NO. 20  
SEQ. ID. NO. 21

SEQ. ID. NO. 18  
SEQ. ID. NO. 19 A C A T G T G G C T C C C C C T G A A T T T T G C  
SEQ. ID. NO. 20  
SEQ. ID. NO. 21

Figure 5q

SEQ. ID. NO. 18  
SEQ. ID. NO. 19 T T C C T T T G G G G A G C T C A T T C T T T C G  
SEQ. ID. NO. 20  
SEQ. ID. NO. 21

SEQ. ID. NO. 18  
SEQ. ID. NO. 19 C C A A G G T C A C A T G C T C C C T T G C C T C  
SEQ. ID. NO. 20  
SEQ. ID. NO. 21

SEQ. ID. NO. 18  
SEQ. ID. NO. 19 T G G C T C C G T G C A  
SEQ. ID. NO. 20  
SEQ. ID. NO. 21

Figure 5r

FIGURE 6a

SEQ. ID. NO. 22 A T G C T G C T G C T G C T G C T G G T G C C T C T C T T C  
SEQ. ID. NO. 23 A T G G G C C C G G G G G G A C C C T G T A C C C C A G T G

SEQ. ID. NO. 22 C T C C G C C C C C T G G G C G C T G G C G G G G C G C A G  
SEQ. ID. NO. 23 G G G T G G C C G C T G C C T C T T C T G C T G G T G A T G

SEQ. ID. NO. 22 A C C C C C A A C G C C A C C T C G G A A G G T T G C C A G  
SEQ. ID. NO. 23 G C G G C T G G G G T G G C T C C G G T G T G G G C C T C T

SEQ. ID. NO. 22 A T T A T A C A T C C G C C C T G G G A A G G T G G C A T C  
SEQ. ID. NO. 23 C A C T C C C C T C A T C T C C C G C G G C C T C A C C C G

SEQ. ID. NO. 22 A G G T A C C G T G G C T T G A C T C G C G A C C A G G T G  
SEQ. ID. NO. 23 A G G G T C C C C C C G C A C C C C T C C T C A G A A C G G

SEQ. ID. NO. 22 A A G G C C A T C A A C T T C C T G C C T G T G G A C T A T  
SEQ. ID. NO. 23 C G T G C A G T A T A C A T C G G G G C G C T G T T T C C C

SEQ. ID. NO. 22 G A G A T C G A A T A T G T G T G C C G A G G G G A G C G C  
SEQ. ID. NO. 23 A T G A G C G G G G G C T G G C C G G G G G G C C A G G C C

SEQ. ID. NO. 22 G A G G T G G T G G G G C C C A A G G T G C G C A A A T G C  
SEQ. ID. NO. 23 T G C C A G C C C G C G G T G G A G A T G G C G C T G G A G

SEQ. ID. NO. 22 C T G G C C A A C G G C T C C T G G A C G G A T A T G G A C  
SEQ. ID. NO. 23 G A C G T T A A C A G C C G C A G A G A C A T C C T G C C G

SEQ. ID. NO. 22 A C A C C C A G C C G C T G T G T C C G A A T C T G C T C C  
SEQ. ID. NO. 23 G A C T A C G A G C T C A A G C T T A T C C A C C A C G A C

SEQ. ID. NO. 22 A A G T C T T A T T T G A C C C T G G A A A A T G G G A A G  
SEQ. ID. NO. 23 A G C A A G T G T G A C C C A G G G C A A G C C A C C A A G

SEQ. ID. NO. 22 G T T T T C C T G A C G G G T G G G G A C C T C C C A G C T  
SEQ. ID. NO. 23 T A C T T G T A C G A A C T A C T C T A C A A T G A C C C C

FIGURE 6b

SEQ. ID. NO. 22 C T G G A T G G A G C C C G G G T G G A G T T C C G A T G T  
SEQ. ID. NO. 23 A T C A A G A T C A T T C T C A T G C C T G G C T G T A G T

SEQ. ID. NO. 22 G A C C C C G A C T T C C A T C T G G T G G G C A G C T C C  
SEQ. ID. NO. 23 T C T G T C T C C A C A C T T G T A G C T G A G G C T G C C

SEQ. ID. NO. 22 C G G A G C G T C T G T A G T C A G G G C C A G T G G A G C  
SEQ. ID. NO. 23 C G G A T G T G G A A C C T T A T T G T G C T C T C A T A T

SEQ. ID. NO. 22 A C C C C C A A G C C C C A C T G C C A G G T G A A T C G A  
SEQ. ID. NO. 23 G G C T C C A G T T C A C C A G C C T T G T C A A A C C G A

SEQ. ID. NO. 22 A C G C C A C A C T C A G A A C G G C G T G C A G T A T A C  
SEQ. ID. NO. 23 C A G C G G T T T C C C A C G T T C T T C C G G A C G C A T

SEQ. ID. NO. 22 A T C G G G G C G C T G T T T C C C A T G A G C G G G G G C  
SEQ. ID. NO. 23 C C A T C C G C C A C A C T C C A C A A T C C C A C C C G G

SEQ. ID. NO. 22 T G G C C G G G G G G C C A G G C C T G C C A G C C C G C G  
SEQ. ID. NO. 23 G T G A A A C T C T T C G A A A A G T G G G G C T G G A A G

SEQ. ID. NO. 22 G T G G A G A T G G C G C T G G A G G A C G T T A A C A G C  
SEQ. ID. NO. 23 A A G A T C G C T A C C A T C C A A C A G A C C A C C G A G

SEQ. ID. NO. 22 C G C A G A G A C A T C C T G C C G G A C T A C G A G C T C  
SEQ. ID. NO. 23 G T C T T C A C C T C A A C G C T G G A T G A C C T G G A G

SEQ. ID. NO. 22 A A G C T T A T C C A C C A C G A C A G C A A G T G T G A C  
SEQ. ID. NO. 23 G A G C G A G T G A A A G A G G C T G G G A T C G A G A T C

SEQ. ID. NO. 22 C C A G G G C A A G C C A C C A A G T A C T T G T A C G A A  
SEQ. ID. NO. 23 A C T T T C C G A C A G A G T T T C T T C T C G G A T C C A

SEQ. ID. NO. 22 C T A C T C T A C A A T G A C C C C A T C A A G A T C A T T  
SEQ. ID. NO. 23 G C T G T G C C T G T T A A A A A C C T G A A G C G T C A A

SEQ. ID. NO. 22 C T C A T G C C T G G C T G T A G T T C T G T C T C C A C A  
SEQ. ID. NO. 23 G A T G C T C G A A T C A T C G T G G G A C T T T T C T A T

FIGURE 6c

SEQ. ID. NO. 22 C T T G T A G C T G A G G C T G C C C G G A T G T G G A A C  
SEQ. ID. NO. 23 G A G A C G G A A G C C C G G A A A G T T T T T T G T G A G

SEQ. ID. NO. 22 C T T A T T G T G C T C T C A T A T G G C T C C A G T T C A  
SEQ. ID. NO. 23 G T C T A T A A G G A A A G G C T C T T T G G G A A G A A G

SEQ. ID. NO. 22 C C A G C C T T G T C A A A C C G A C A G C G G T T T C C C  
SEQ. ID. NO. 23 T A C G T C T G G T T C C T C A T C G G G T G G T A T G C T

SEQ. ID. NO. 22 A C G T T C T T C C G G A C G C A T C C A T C C G C C A C A  
SEQ. ID. NO. 23 G A C A A C T G G T T C A A G A C C T A T G A C C C G T C A

SEQ. ID. NO. 22 C T C C A C A A T C C C A C C C G G G T G A A A C T C T T C  
SEQ. ID. NO. 23 A T C A A T T G T A C A G T G G A A G A A A T G A C C G A G

SEQ. ID. NO. 22 G A A A A G T G G G G C T G G A A G A A G A T C G C T A C C  
SEQ. ID. NO. 23 G C G G T G G A G G G C C A C A T C A C C A C G G A G A T T

SEQ. ID. NO. 22 A T C C A A C A G A C C A C C G A G G T C T T C A C C T C A  
SEQ. ID. NO. 23 G T C A T G C T G A A C C C T G C C A A C A C C C G A A G C

SEQ. ID. NO. 22 A C G C T G G A T G A C C T G G A G G A G C G A G T G A A A  
SEQ. ID. NO. 23 A T T T C C A A C A T G A C G T C A C A G G A A T T T G T G

SEQ. ID. NO. 22 G A G G C T G G G A T C G A G A T C A C T T T C C G A C A G  
SEQ. ID. NO. 23 G A G A A A C T A A C C A A G C G G C T G A A A A G A C A C

SEQ. ID. NO. 22 A G T T T C T T C T C G G A T C C A G C T G T G C C T G T T  
SEQ. ID. NO. 23 C C C G A G G A G A C T G G A G G C T T C C A G G A G G C A

SEQ. ID. NO. 22 A A A A A C C T G A A G C G T C A A G A T G C T C G A A T C  
SEQ. ID. NO. 23 C C A C T G G C C T A T G A T G C T A T C T G G G C C T T G

SEQ. ID. NO. 22 A T C G T G G G A C T T T T C T A T G A G A C G G A A G C C  
SEQ. ID. NO. 23 G C T T T G G C C T T G A A C A A G A C G T C T G G A G G A

SEQ. ID. NO. 22 C G G A A A G T T T T T G T G A G G T C T A T A A G G A A  
SEQ. ID. NO. 23 G G T G G T C G T T C C G G C G T G C G C C T G G A G G A C



FIGURE 6d

SEQ. ID. NO. 22 A G G C T C T T T G G G A A G A A G T A C G T C T G G T T C  
SEQ. ID. NO. 23 T T T A A C T A C A A C A A C C A G A C C A T T A C A G A C

SEQ. ID. NO. 22 C T C A T C G G G T G G T A T G C T G A C A A C T G G T T C  
SEQ. ID. NO. 23 C A G A T C T A C C G G G C C A T G A A C T C C T C C T C C

SEQ. ID. NO. 22 A A G A C C T A T G A C C C G T C A A T C A A T T G T A C A  
SEQ. ID. NO. 23 T T T G A G G G C G T T T C T G G C C A T G T G G T C T T T

SEQ. ID. NO. 22 G T G G A A G A A A T G A C C G A G G C G G T G G A G G G C  
SEQ. ID. NO. 23 G A T G C C A G C G G C T C C C G G A T G G C A T G G A C A

SEQ. ID. NO. 22 C A C A T C A C C A C G G A G A T T G T C A T G C T G A A C  
SEQ. ID. NO. 23 C T T A T C G A G C A G C T A C A G G G C G G C A G C T A C

SEQ. ID. NO. 22 C C T G C C A A C A C C C G A A G C A T T T C C A A C A T G  
SEQ. ID. NO. 23 A A G A A G A T C G G C T A C T A C G A C A G C A C C A A G

SEQ. ID. NO. 22 A C G T C A C A G G A A T T T G T G G A G A A A C T A A C C  
SEQ. ID. NO. 23 G A T G A T C T T T C C T G G T C C A A A A C G G A C A A G

SEQ. ID. NO. 22 A A G C G G C T G A A A A G A C A C C C C G A G G A G A C T  
SEQ. ID. NO. 23 T G G A T T G G A G G G T C T C C C C C A G C T G A C C A G

SEQ. ID. NO. 22 G G A G G C T T C C A G G A G G C A C C A C T G G C C T A T  
SEQ. ID. NO. 23 A C C T T G G T C A T C A A G A C A T T C C G T T T C C T G

SEQ. ID. NO. 22 G A T G C T A T C T G G G C C T T G G C T T T G G C C T T G  
SEQ. ID. NO. 23 T C T C A G A A A C T C T T T A T C T C C G T C T C A G T T

SEQ. ID. NO. 22 A A C A A G A C G T C T G G A G G A G G T G G T C G T T C C  
SEQ. ID. NO. 23 C T C T C C A G C C T G G G C A T T G T T C T T G C T G T T

SEQ. ID. NO. 22 G G C G T G C G C C T G G A G G A C T T T A A C T A C A A C  
SEQ. ID. NO. 23 G T C T G T C T G T C C T T T A A C A T C T A C A A C T C C

SEQ. ID. NO. 22 A A C C A G A C C A T T A C A G A C C A G A T C T A C C G G  
SEQ. ID. NO. 23 C A C G T T C G T T A T A T C C A G A A C T C C C A G C C C

FIGURE 6e

SEQ. ID. NO. 22 G C C A T G A A C T C C T C C T C C T T T G A G G G C G T T  
SEQ. ID. NO. 23 A A C C T G A A C A A T C T G A C T G C T G T G G G C T G C

SEQ. ID. NO. 22 T C T G G C C A T G T G G T C T T T G A T G C C A G C G G C  
SEQ. ID. NO. 23 T C A C T G G C A C T G G C T G C T G T C T T C C C T C T C

SEQ. ID. NO. 22 T C C C G G A T G G C A T G G A C A C T T A T C G A G C A G  
SEQ. ID. NO. 23 G G G C T G G A T G G T T A C C A C A T A G G G A G A A G C

SEQ. ID. NO. 22 C T A C A G G G C G G C A G C T A C A A G A A G A T C G G C  
SEQ. ID. NO. 23 C A G T T C C C G T T T G T C T G C C A G G C C C G C C T T

SEQ. ID. NO. 22 T A C T A C G A C A G C A C C A A G G A T G A T C T T T C C  
SEQ. ID. NO. 23 T G G C T C T T G G G C T T G G G C T T T A G T C T G G G C

SEQ. ID. NO. 22 T G G T C C A A A A C G G A C A A G T G G A T T G G A G G G  
SEQ. ID. NO. 23 T A T G G C T C T A T G T T C A C C A A G A T C T G G T G G

SEQ. ID. NO. 22 T C T C C C C A G C T G A C C A G A C C T T G G T C A T C  
SEQ. ID. NO. 23 G T C C A C A C A G T C T T C A C G A A G A A G G A G G A G

SEQ. ID. NO. 22 A A G A C A T T C C G T T T C C T G T C T C A G A A A C T C  
SEQ. ID. NO. 23 A A G A A G G A G T G G A G G A A G A C C C T A G A G C C C

SEQ. ID. NO. 22 T T T A T C T C C G T C T C A G T T C T C T C C A G C C T G  
SEQ. ID. NO. 23 T G G A A A C T C T A T G C C A C T G T G G G C C T G C T G

SEQ. ID. NO. 22 G G C A T T G T T C T T G C T G T T G T C T G T C T G T C C  
SEQ. ID. NO. 23 G T G G G C A T G G A T G T C C T G A C T C T T G C C A T C

SEQ. ID. NO. 22 T T T A A C A T C T A C A A C T C C C A C G T T C G T T A T  
SEQ. ID. NO. 23 T G G C A G A T T G T G G A C C C C T T G C A C C G A A C C

SEQ. ID. NO. 22 A T C C A G A A C T C C C A G C C C A A C C T G A A C A A T  
SEQ. ID. NO. 23 A T T G A G A C T T T T G C C A A G G A G G A A C C A A A G

SEQ. ID. NO. 22 C T G A C T G C T G T G G G C T G C T C A C T G G C A C T G  
SEQ. ID. NO. 23 G A A G A C A T C G A T G T C T C C A T T C T G C C C C A G

FIGURE 6f

SEQ. ID. NO. 22 G C T G C T G T C T T C C C T C T C G G G C T G G A T G G T  
SEQ. ID. NO. 23 T T G G A G C A C T G C A G C T C C A A G A A G A T G A A T

SEQ. ID. NO. 22 T A C C A C A T A G G G A G A A G C C A G T T C C C G T T T  
SEQ. ID. NO. 23 A C G T G G C T T G G C A T T T T C T A T G G T T A C A A G

SEQ. ID. NO. 22 G T C T G C C A G G C C C G C C T T T G G C T C T T G G G C  
SEQ. ID. NO. 23 G G G C T G C T G C T G C T G C T G G G A A T C T T T C T T

SEQ. ID. NO. 22 T T G G G C T T T A G T C T G G G C T A T G G C T C T A T G  
SEQ. ID. NO. 23 G C T T A C G A A A C C A A G A G C G T G T C C A C T G A A

SEQ. ID. NO. 22 T T C A C C A A G A T C T G G T G G G T C C A C A C A G T C  
SEQ. ID. NO. 23 A A G A T C A A T G A C C A C A G G G C C G T G G G C A T G

SEQ. ID. NO. 22 T T C A C G A A G A A G G A G G A G A A G A A G G A G T G G  
SEQ. ID. NO. 23 G C T A T C T A C A A T G T C G C G G T C C T G T G T C T C

SEQ. ID. NO. 22 A G G A A G A C C C T A G A G C C C T G G A A A C T C T A T  
SEQ. ID. NO. 23 A T C A C T G C T C C T G T G A C C A T G A T C C T T T C C

SEQ. ID. NO. 22 G C C A C T G T G G G C C T G C T G G T G G G C A T G G A T  
SEQ. ID. NO. 23 A G T C A G C A G G A C G C A G C C T T T G C C T T T G C C

SEQ. ID. NO. 22 G T C C T G A C T C T T G C C A T C T G G C A G A T T G T G  
SEQ. ID. NO. 23 T C T C T G G C C A T C G T G T T C T C T T C C T A C A T C

SEQ. ID. NO. 22 G A C C C C T T G C A C C G A A C C A T T G A G A C T T T T  
SEQ. ID. NO. 23 A C T C T G G T T G T G C T C T T T G T G C C C A A G A T G

SEQ. ID. NO. 22 G C C A A G G A G G A A C C A A A G G A A G A C A T C G A T  
SEQ. ID. NO. 23 C G C A G G C T G A T C A C C C G A G G G G A A T G G C A G

SEQ. ID. NO. 22 G T C T C C A T T C T G C C C C A G T T G G A G C A C T G C  
SEQ. ID. NO. 23 T C T G A A A C G C A G G A C A C C A T G A A A A C A G G A

SEQ. ID. NO. 22 A G C T C C A A G A A G A T G A A T A C G T G G C T T G G C  
SEQ. ID. NO. 23 T C A T C C A C C A A C A A C A A C G A G G A A G A G A A G

FIGURE 6c

SEQ. ID. NO. 22 A T T T T C T A T G G T T A C A A G G G G C T G C T G C T G  
SEQ. ID. NO. 23 T C C C G A C T G T T G G A G A A G G A A A A C C G A G A A

SEQ. ID. NO. 22 C T G C T G G G A A T C T T T C T T G C T T A C G A A A C C  
SEQ. ID. NO. 23 C T G G A A A A G A T C A T C G C T G A G A A A G A G G A G

SEQ. ID. NO. 22 A A G A G C G T G T C C A C T G A A A A G A T C A A T G A C  
SEQ. ID. NO. 23 C G C G T C T C T G A A C T G C G C C A T C A G C T C C A G

SEQ. ID. NO. 22 C A C A G G G C C G T G G G C A T G G C T A T C T A C A A T  
SEQ. ID. NO. 23 T C T C G G C A G C A A C T C C G C T C A C G G C G C C A C

SEQ. ID. NO. 22 G T C G C G G T C C T G T G T C T C A T C A C T G C T C C T  
SEQ. ID. NO. 23 C C C C C A A C A C C C C C A G A T C C C T C T G G G G G C

SEQ. ID. NO. 22 G T G A C C A T G A T C C T T T C C A G T C A G C A G G A C  
SEQ. ID. NO. 23 C T T C C C A G G G G A C C C T C T G A G C C C C C T G A C

SEQ. ID. NO. 22 G C A G C C T T T G C C T T T G C C T C T C T G G C C A T C  
SEQ. ID. NO. 23 C G G C T T A G C T G T G A T G G G A G T C G A G T A C A T

SEQ. ID. NO. 22 G T G T T C T C T T C C T A C A T C A C T C T G G T T G T G  
SEQ. ID. NO. 23 T T G C T T T A C A A G

SEQ. ID. NO. 22 C T C T T T G T G C C C A A G A T G C G C A G G C T G A T C  
SEQ. ID. NO. 23

SEQ. ID. NO. 22 A C C C G A G G G G A A T G G C A G T C T G A A A C G C A G  
SEQ. ID. NO. 23

SEQ. ID. NO. 22 G A C A C C A T G A A A A C A G G A T C A T C C A C C A A C  
SEQ. ID. NO. 23

SEQ. ID. NO. 22 A A C A A C G A G G A A G A G A A G T C C C G A C T G T T G  
SEQ. ID. NO. 23

SEQ. ID. NO. 22 G A G A A G G A A A A C C G A G A A C T G G A A A A G A T C  
SEQ. ID. NO. 23

FIGURE 61.

SEQ. ID. NO. 22 A T C G C T G A G A A A G A G G A G C G C G T C T C T G A A  
SEQ. ID. NO. 23

SEQ. ID. NO. 22 C T G C G C C A T C A G C T C C A G T C T C G G C A G C A A  
SEQ. ID. NO. 23

SEQ. ID. NO. 22 C T C C G C T C A C G G C G C C A C C C C C C A A C A C C C  
SEQ. ID. NO. 23

SEQ. ID. NO. 22 C C A G A T C C C T C T G G G G G C C T T C C C A G G G G A  
SEQ. ID. NO. 23

SEQ. ID. NO. 22 C C C T C T G A G C C C C C T G A C C G G C T T A G C T G T  
SEQ. ID. NO. 23

SEQ. ID. NO. 22 G A T G G G A G T C G A G T A C A T T T G C T T T A C A A G  
SEQ. ID. NO. 23

FIGURE 7c

SEQ. ID. NO. 24 M L L L L L V P L F L R P L G A G G A Q T P N A T S E G C Q  
 SEQ. ID. NO. 25 M G P G G P C T P V G W P L P L L L V M A A G V A P V W A S

SEQ. ID. NO. 24 I I H P P W E G G I R Y R G L T R D Q V K A I N F L P V D Y  
 SEQ. ID. NO. 25 H S P H L P R P H P R V P P H P S S E R R A V Y I G A L F P

SEQ. ID. NO. 24 E I E Y V C R G E R E V V G P K V R K C L A N G S W T D M D  
 SEQ. ID. NO. 25 M S G G W P G G Q A C Q P A V E M A L E D V N S R R D I L P

SEQ. ID. NO. 24 T P S R C V R I C S K S Y L T L E N G K V F L T G G D L P A  
 SEQ. ID. NO. 25 D Y E L K L I H H D S K C D P G Q A T K Y L Y E L L Y N D P

SEQ. ID. NO. 24 L D G A R V E F R C D P D F H L V G S S R S V C S Q G Q W S  
 SEQ. ID. NO. 25 I K I I L M P G C S S V S T L V A E A A R M W N L I V L S Y

SEQ. ID. NO. 24 T P K P H C Q V N R T P H S E R R A V Y I G A L F P M S G G  
 SEQ. ID. NO. 25 G S S S P A L S N R Q R F P T F F R T H P S A T L H N P T R

SEQ. ID. NO. 24 W P G G Q A C Q P A V E M A L E D V N S R R D I L P D Y E L  
 SEQ. ID. NO. 25 V K L F E K W G W K K I A T I Q Q T T E V F T S T L D D L E

SEQ. ID. NO. 24 K L I H H D S K C D P G Q A T K Y L Y E L L Y N D P I K I I  
 SEQ. ID. NO. 25 E R V K E A G I E I T F R Q S F F S D P A V P V K N L K R Q

SEQ. ID. NO. 24 L M P G C S S V S T L V A E A A R M W N L I V L S Y G S S S  
 SEQ. ID. NO. 25 D A R I I V G L F Y E T E A R K V F C E V Y K E R L F G K K

SEQ. ID. NO. 24 P A L S N R Q R F P T F F R T H P S A T L H N P T R V K L F  
 SEQ. ID. NO. 25 Y V W F L I G W Y A D N W F K T Y D P S I N C T V E E M T E

SEQ. ID. NO. 24 E K W G W K K I A T I Q Q T T E V F T S T L D D L E E R V K  
 SEQ. ID. NO. 25 A V E G H I T T E I V M L N P A N T R S I S N M T S Q E F V

SEQ. ID. NO. 24 E A G I E I T F R Q S F F S D P A V P V K N L K R Q D A R I  
 SEQ. ID. NO. 25 E K L T K R L K R H P E E T G G F Q E A P L A Y D A I W A L

FIGURE 71

SEQ. ID. NO. 24 I V G L F Y E T E A R K V F C E V Y K E R L F G K K Y V W F  
SEQ. ID. NO. 25 A L A L N K T S G G G G R S G V R L E D F N Y N N Q T I T D

SEQ. ID. NO. 24 L I G W Y A D N W F K T Y D P S I N C T V E E M T E A V E G  
SEQ. ID. NO. 25 Q I Y R A M N S S S F E G V S G H V V F D A S G S R M A W T

SEQ. ID. NO. 24 H I T T E I V M L N P A N T R S I S N M T S Q E F V E K L T  
SEQ. ID. NO. 25 L I E Q L Q G G S Y K K I G Y Y D S T K D D L S W S K T D K

SEQ. ID. NO. 24 K R L K R H P E E T G G F Q E A P L A Y D A I W A L A L A L  
SEQ. ID. NO. 25 W I G G S P P A D Q I L V I K T F R F L S Q K L F I S V S V

SEQ. ID. NO. 24 N K T S G G G G R S G V R L E D F N Y N N Q T I T D Q I Y R  
SEQ. ID. NO. 25 L S S L G I V L A V V C L S F N I Y N S H V R Y I Q N S Q P

SEQ. ID. NO. 24 A M N S S S F E G V S G H V V F D A S G S R M A W T L I E Q  
SEQ. ID. NO. 25 N L N N L T A V G C S L A L A A V F P L G L D G Y H I G R S

SEQ. ID. NO. 24 L Q G G S Y K K I G Y Y D S T K D D L S W S K T D K W I G G  
SEQ. ID. NO. 25 Q F P F V C Q A R L W L L G L G F S L G Y G S M F T K I W W

SEQ. ID. NO. 24 S P P A D Q I L V I K T F R F L S Q K L F I S V S V L S S L  
SEQ. ID. NO. 25 V H T V F T K K E E K K E W R K T L E P W K L Y A T V G L L

SEQ. ID. NO. 24 G I V L A V V C L S F N I Y N S H V R Y I Q N S Q P N L N N  
SEQ. ID. NO. 25 V G M D V L T L A I W Q I V D P L H R T I E T F A K E E P K

SEQ. ID. NO. 24 L T A V G C S L A L A A V F P L G L D G Y H I G R S Q F P F  
SEQ. ID. NO. 25 E D I D V S I L P Q L E H C S S K K M N T W L G I F Y G Y K

SEQ. ID. NO. 24 V C Q A R L W L L G L G F S L G Y G S M F T K I W W V H T V  
SEQ. ID. NO. 25 G L L L L L G I F L A Y E T K S V S T E K I N D H R A V G M

SEQ. ID. NO. 24 F T K K E E K K E W R K T L E P W K L Y A T V G L L V G M D  
SEQ. ID. NO. 25 A I Y N V A V L C L I T A P V T M I L S S Q Q D A A F A F A

SEQ. ID. NO. 24 V L T L A I W Q I V D P L H R T I E T F A K E E P K E D I D  
SEQ. ID. NO. 25 S L A I V F S S Y I T L V V L F V P K M R R L I T R G E W Q

FIGURE 7c

SEQ. ID. NO. 24 V S I L P Q L E H C S S K K M N T W L G I F Y G Y K G L L L  
SEQ. ID. NO. 25 S E T Q D T M K T G S S T N N N E E E K S R L L E K E N R E

SEQ. ID. NO. 24 L L G I F L A Y E T K S V S T E K I N D H R A V G M A I Y N  
SEQ. ID. NO. 25 L E K I I A E K E E R V S E L R H Q L Q S R Q Q L R S R R H

SEQ. ID. NO. 24 V A V L C L I T A P V T M I L S S Q Q D A A F A F A S L A I  
SEQ. ID. NO. 25 P P T P P D P S G G L P R G P S E P P D R L S C D G S R V H

SEQ. ID. NO. 24 V F S S Y I T L V V L F V P K M R R L I T R G E W Q S E T Q  
SEQ. ID. NO. 25 L L Y K

SEQ. ID. NO. 24 D T M K T G S S T N N N E E E K S R L L E K E N R E L E K I  
SEQ. ID. NO. 25

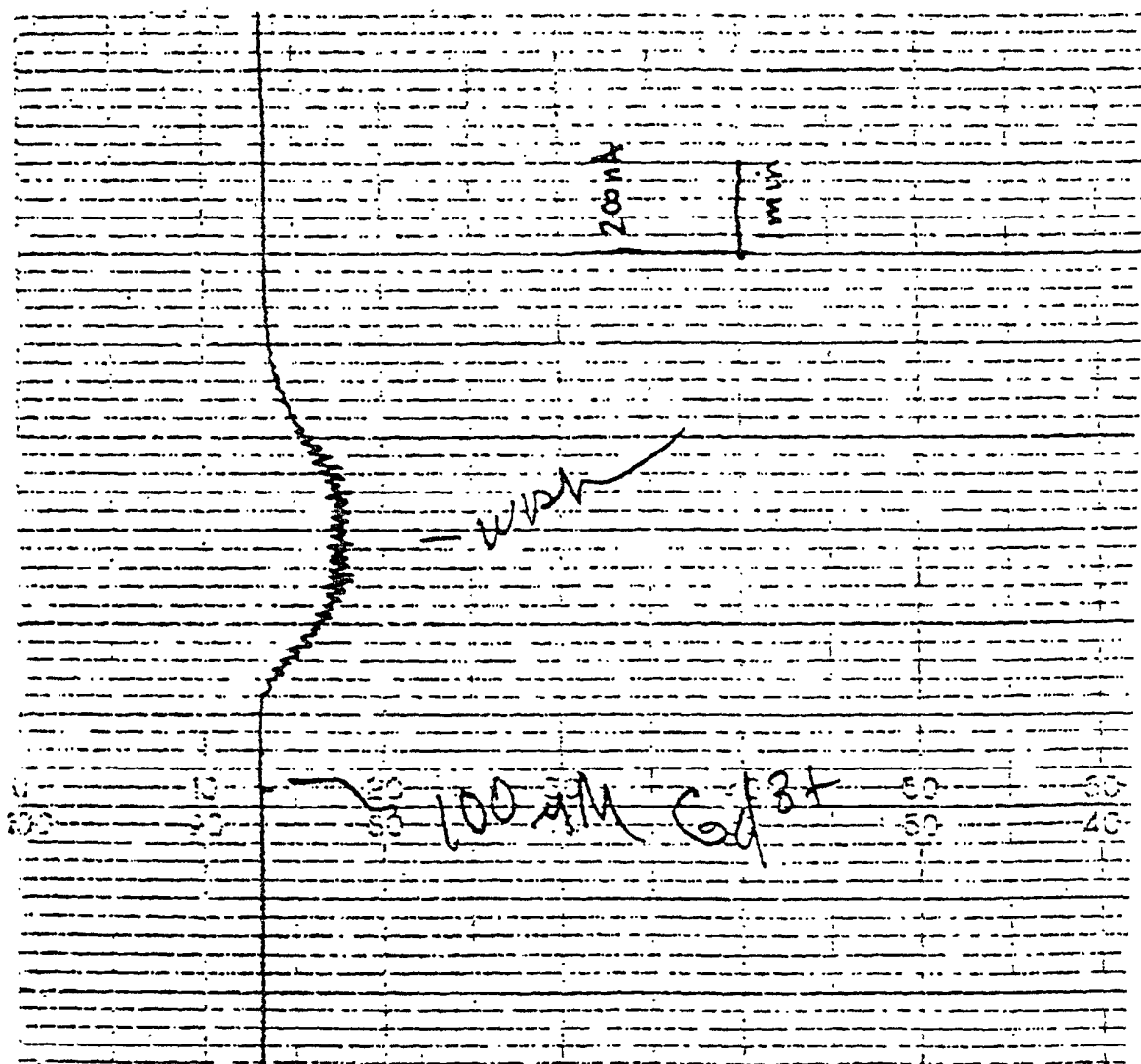
SEQ. ID. NO. 24 I A E K E E R V S E L R H Q L Q S R Q Q L R S R R H P P T P  
SEQ. ID. NO. 25

SEQ. ID. NO. 24 P D P S G G L P R G P S E P P D R L S C D G S R V H L L Y K  
SEQ. ID. NO. 25



FIGURE 8

CHART NO. NZ 200 0



# ClustalW Formatted Alignments

SEQ. ID. NO. 38 A T G G T A T G C G A G G G A A A G C G A T C A G  
 SEQ. ID. NO. 34 A T G G G A T C G C T G C T T G C G C T C C C G G  
 SEQ. ID. NO. 30 A T G G C A T T T T A T A G C T G C T G C T G G G  
 SEQ. ID. NO. 26 A T G G G A T C G C T G C T T G C G C T C C T G G

SEQ. ID. NO. 38 C C T C T T G C C C T T G T T T C T T C C T C T T  
 SEQ. ID. NO. 34 C A C T G C T G C T G C T G T G G G G T G C T G T  
 SEQ. ID. NO. 30 T C C T C T T G G C A C T C A C C T G G C A C A C  
 SEQ. ID. NO. 26 C A C T G C T G C C G C T G T G G G G T G C T G T

SEQ. ID. NO. 38 G A C C G C C A A G T T C T A C T G G A T C C T C  
 SEQ. ID. NO. 34 G G C T G A G G G C C C A G C C A A G A A G G T G  
 SEQ. ID. NO. 30 C T C T G C C T A C G G G C C A G A C C A G C G A  
 SEQ. ID. NO. 26 G G C T G A G G G C C C A G C C A A G A A G G T G

SEQ. ID. NO. 38 A C A A T G A T G C A A A G A A C T C A C A G C C  
 SEQ. ID. NO. 34 C T G A C C C T G G A G G G A G A C T T G G T G C  
 SEQ. ID. NO. 30 G C C C A A A A G A A A G G G G G A C A T T A T C C  
 SEQ. ID. NO. 26 C T G A C C C T G G A G G G A G A C T T G G T G C

SEQ. ID. NO. 38 A G G A G T A T G C C C A T T C C A T A C G G G T  
 SEQ. ID. NO. 34 T G G G T G G G C T G T T C C C A G T G C A C C A  
 SEQ. ID. NO. 30 T T G G G G G G C T C T T T C C T A T T C A T T T  
 SEQ. ID. NO. 26 T G G G T G G G C T G T T C C C A G T G C A C C A

SEQ. ID. NO. 38 G G A T G G G G A C A T T A T T T T G G G G G G T  
 SEQ. ID. NO. 34 G A A G G G C G G C C C A G C A G A G G A C T G T  
 SEQ. ID. NO. 30 T G G A G T A G C A G C T A A A G A T C A A G A T  
 SEQ. ID. NO. 26 G A A G G G C G G C C C A G C A G A G G A C T G T

SEQ. ID. NO. 38 C T C T T C C C T G T C C A C G C A A A G G G A G  
 SEQ. ID. NO. 34 G G T C C T G T C A A T G A G C A C C G T G G C A  
 SEQ. ID. NO. 30 C T C A A A T C A A A G G C C G G A G T C T G T G G  
 SEQ. ID. NO. 26 G G T C C T G T C A A T G A G C A C C G T G G C A

SEQ. ID. NO. 38 A G A G A G G G G T G C C T T G T G G G G A G C T  
 SEQ. ID. NO. 34 T C C A G C G C C T G G A G G C C A T G C T T T T  
 SEQ. ID. NO. 30 A A T G T A T C A G G T A T A A T T T C C G T G G  
 SEQ. ID. NO. 26 T C C A G C G C C T G G A G G C C A T G C T T T T

Figure 9a

SEQ. ID. NO. 38 G A A G A A G G A A A A G G G G A T T C A C A G A  
 SEQ. ID. NO. 34 T G C A C T G G A C C G C A T C A A C C G T G A C  
 SEQ. ID. NO. 30 G T T T C G C T G G T T A C A G G C T A T G A T A  
 SEQ. ID. NO. 26 T G C A C T G G A C C G C A T C A A C C G T G A C

SEQ. ID. NO. 38 C T G G A G G C C A T G C T T T A T G C A A T T G  
 SEQ. ID. NO. 34 C C G C A C C T G C T G C C T G G C G T G C G C C  
 SEQ. ID. NO. 30 T T T G C C A T A G A G G A G A T A A A C A G C A  
 SEQ. ID. NO. 26 C C G C A C C T G C T G C C T G G C G T G C G C C

SEQ. ID. NO. 38 A C C A G A T T A A C A A G G A C C C T G A T C T  
 SEQ. ID. NO. 34 T G G G T G C A C A C A T C C T C G A C A G T T G  
 SEQ. ID. NO. 30 G C C C A G C C C T T C T T C C C A A C T T G A C  
 SEQ. ID. NO. 26 T G G G T G C A C A C A T C C T C G A C A G T T G

SEQ. ID. NO. 38 C C T T T C C A A C A T C A C T C T G G G T G T C  
 SEQ. ID. NO. 34 C T C C A A G G A C A C A C A T G C G C T G G A G  
 SEQ. ID. NO. 30 G C T G G G A T A C A G G A T A T T T G A C A C T  
 SEQ. ID. NO. 26 C T C C A A G G A C A C A C A T G C G C T G G A G

SEQ. ID. NO. 38 C G C A T C C T C G A C A C G T G C T C T A G G G  
 SEQ. ID. NO. 34 C A G G C A C T G G A C T T T G T G C G T G C C T  
 SEQ. ID. NO. 30 T G C A A C A C C G T T T C T A A G G C C T T G G  
 SEQ. ID. NO. 26 C A G G C A C T G G A C T T T G T G C G T G C C T

SEQ. ID. NO. 38 A C A C C T A T G C T T T T G G A G C A G T C T C T  
 SEQ. ID. NO. 34 C A C T C A G C C G T G G T G C T G A T G G C T C  
 SEQ. ID. NO. 30 A A G C C A C C C T G A G T T T T G T T G C T C A  
 SEQ. ID. NO. 26 C A C T C A G C C G T G G T G C T G A T G G A T C

SEQ. ID. NO. 38 A A C A T T C G T G C A G G C A T T A A T A G A G  
 SEQ. ID. NO. 34 A C G C C A C A T C T G C C C C G A C G G C T C T  
 SEQ. ID. NO. 30 A A A C A A A A T T G A T T C T T T G A A C C T T  
 SEQ. ID. NO. 26 A C G C C A C A T C T G C C C C G A C G G C T C T

SEQ. ID. NO. 38 A A A G A T G C T T C G G A T G T G A A G T G T G  
 SEQ. ID. NO. 34 T A T G C G A C C C A T G G T G A T G C T C C C A  
 SEQ. ID. NO. 30 G A T G A G T T C T G C A A C T G C T C A G A G C  
 SEQ. ID. NO. 26 T A T G C G A C C C A T G G T G A T G C T C C C A

Figure 9b

SEQ. ID. NO. 38 C T A A T G G A G A T C C A C C C A T T T T C A C  
 SEQ. ID. NO. 34 C T G C C A T C A C T G G T G T T A T T G G C G G  
 SEQ. ID. NO. 30 A C A T T C C C T C T A C G A T T G C T G T G G T  
 SEQ. ID. NO. 26 C T G C C A T C A C T G G T G T T A T T G G C G G

SEQ. ID. NO. 38 C A A G C C C G A C A A G A T T T C T G G C G T C  
 SEQ. ID. NO. 34 T T C C T A C A G T G A T G T C T C C A T C C A G  
 SEQ. ID. NO. 30 G G G A G C A A C T G G C T C A G G C G T C T C C  
 SEQ. ID. NO. 26 T T C C T A C A G T G A T G T C T C C A T C C A G

SEQ. ID. NO. 38 A T A G G T G C T G C A G C A A G C T C C G T G T  
 SEQ. ID. NO. 34 G T G G C C A A C C T C T T G A G G C T A T T T C  
 SEQ. ID. NO. 30 A C G G C A G T G G C A A A T C T G C T G G G G C  
 SEQ. ID. NO. 26 G T G G C C A A C C T C T T G A G G C T A T T T C

SEQ. ID. NO. 38 C C A T C A T G G T T G C T A A C A T T T T A A G  
 SEQ. ID. NO. 34 A G A T C C C A C A G A T T A G C T A C G C C T C  
 SEQ. ID. NO. 30 T C T T C T A C A T T C C C C A G G T C A G T T A  
 SEQ. ID. NO. 26 A G A T C C C A C A G A T T A G C T A C G C C T C

SEQ. ID. NO. 38 A C T T T T T A A G A T A C C T C A A A T C A G C  
 SEQ. ID. NO. 34 T A C C A G T G C C A A G C T G A G T G A C A A G  
 SEQ. ID. NO. 30 T G C C T C C T C C A G C A G A C T C C T C A G C  
 SEQ. ID. NO. 26 T A C C A G T G C C A A G C T G A G T G A C A A G

SEQ. ID. NO. 38 T A T G C A T C C A C A G C C C C A G A G C T A A  
 SEQ. ID. NO. 34 T C C C G C T A T G A C T A C T T T G C C C G C A  
 SEQ. ID. NO. 30 A A C A A G A A T C A A T T C A A G T C T T T C C  
 SEQ. ID. NO. 26 T C C C G C T A T G A C T A C T T T G C C C G C A

SEQ. ID. NO. 38 G T G A T A A C A C C A G G T A T G A C T T T T T  
 SEQ. ID. NO. 34 C A G T G C C T C C T G A C T T C T T C C A A G C  
 SEQ. ID. NO. 30 T C C G A A C C A T C C C C A A T G A T G A G C A  
 SEQ. ID. NO. 26 C A G T G C C T C C T G A C T T C T T C C A A G C

SEQ. ID. NO. 38 C T C T C G A G T G G T T C C G C C T G A C T C C  
 SEQ. ID. NO. 34 C A A G G C C A T G G C T G A G A T T C T C C G C  
 SEQ. ID. NO. 30 C C A G G C C A C T G C C A T G G C A G A C A T C  
 SEQ. ID. NO. 26 C A A G G C C A T G G C T G A G A T T C T C C G C

Figure 9c

SEQ. ID. NO. 38 T A C C A A G C C C A A G C C A T G G T G G A C A  
 SEQ. ID. NO. 34 T T C T T C A A C T G G A C C T A T G T G T C C A  
 SEQ. ID. NO. 30 A T C G A G T A T T T C C G C T G G A A C T G G G  
 SEQ. ID. NO. 26 T T C T T C A A C T G G A C C T A T G T G T C C A

SEQ. ID. NO. 38 T C G T G A C A G C A C T G G G A T G G A A T T A  
 SEQ. ID. NO. 34 C T G T G G C G T C T G A G G G C G A C T A T G G  
 SEQ. ID. NO. 30 T G G G C A C A A T T G C A G C T G A T G A C G A  
 SEQ. ID. NO. 26 C T G A G G C C T C T G A G G G C G A C T A T G G

SEQ. ID. NO. 38 T G T T T C G A C A C T G G C T T C T G A G G G G  
 SEQ. ID. NO. 34 C G A G A C A G G C A T T G A G G C C T T T G A G  
 SEQ. ID. NO. 30 C T A T G G G C G G C C G G G A T T G A G A A A  
 SEQ. ID. NO. 26 C G A G A C A G G C A T T G A G G C C T T T G A G

SEQ. ID. NO. 38 A A C T A T G G T G A G A G C G G T G T G G A G G  
 SEQ. ID. NO. 34 C T A G A G G C T C G T G C C C G C A A C A T C T  
 SEQ. ID. NO. 30 T T C C G A G A G G A A G C T G A G G A A A G G G  
 SEQ. ID. NO. 26 C T A G A G G C T C G T G C C C G C A A C A T C T

SEQ. ID. NO. 38 C C T T C A C C C A G A T C T C G A G G G A G A T  
 SEQ. ID. NO. 34 G T G T G G C C A C C T C G G A G A A A G T G G G  
 SEQ. ID. NO. 30 A T A T C T G C A T C G A C T T C A G T G A A C T  
 SEQ. ID. NO. 26 G T G T G G C C A C C T C G G A G A A A G T G G G

SEQ. ID. NO. 38 T G G T G G T G T T T G C A T T G C T C A G T C A  
 SEQ. ID. NO. 34 C C G T G C C A T G A G C C G C G C G G C C T T T  
 SEQ. ID. NO. 30 C A T C T C C C A G T A C T C T G A T G A G G A A  
 SEQ. ID. NO. 26 C C G T G C C A T G A G C C G C G C G G C C T T T

SEQ. ID. NO. 38 C A G A A A A T C C C A C G T G A A C C A A G A C  
 SEQ. ID. NO. 34 G A G G G T G T G G T G C G A G C C C T G C T G C  
 SEQ. ID. NO. 30 G A G A T C C A G C A T G T G G T A G A G G T G A  
 SEQ. ID. NO. 26 G A G G G T G T G G T G C G A G C C C T G C T G C

SEQ. ID. NO. 38 C T G G A G A A T T T G A A A A A T T A T C A A  
 SEQ. ID. NO. 34 A G A A G C C C A G T G C C C G C G T G G C T G T  
 SEQ. ID. NO. 30 T T C A A A A T T C C A C G G C C A A A G T C A T  
 SEQ. ID. NO. 26 A G A A G C C C A G T G C C C G C G T G G C T G T

Figure 9d

SEQ. ID. NO. 38 A C G C C T G C T A G A A A C A C C T A A T G C T  
 SEQ. ID. NO. 34 C C T G T T C A C C C G T T C T G A G G A T G C C  
 SEQ. ID. NO. 30 C G T G G T T T T C T C C A G T G G C C C A G A T  
 SEQ. ID. NO. 26 C C T G T T C A C C C G T T C T G A G G A T G C C

SEQ. ID. NO. 38 C G A G C A G T G A T T A T G T T T G C C A A T G  
 SEQ. ID. NO. 34 C G G G A G C T G C T T G C T G C C A G C C A G C  
 SEQ. ID. NO. 30 C T T G A G C C C C T C A T C A A G G A G A T T G  
 SEQ. ID. NO. 26 C G G G A G C T G C T T G C T G C C A G C C A G C

SEQ. ID. NO. 38 A G G A T G A C A T C A G G A G G A T A T T G G A  
 SEQ. ID. NO. 34 G C C T C A A T G C C A G C T T C A C C T G G G T  
 SEQ. ID. NO. 30 T C C G G C G C A A T A T C A C G G G C A A G A T  
 SEQ. ID. NO. 26 G C C T C A A T G C C A G C T T C A C C T G G G T

SEQ. ID. NO. 38 A G C A G C A A A A A A C T A A A C C A A A G T  
 SEQ. ID. NO. 34 G G C C A G T G A T G G T T G G G G G G C C C T G  
 SEQ. ID. NO. 30 C T G G C T G G C C A G C G A G G C C T G G G C C  
 SEQ. ID. NO. 26 G G C C A G T G A T G G T T G G G G G G C C C T G

SEQ. ID. NO. 38 G G G C A T T T T C T C T G G A T T G G C T C A G  
 SEQ. ID. NO. 34 G A G A G T G T G G T G G C A G G C A G T G A G G  
 SEQ. ID. NO. 30 A G C T C C T C C C T G A T C G C C A T G C C T C  
 SEQ. ID. NO. 26 G A G A G T G T G G T G G C A G G C A G T G A G G

SEQ. ID. NO. 38 A T A G T T G G G G A T C C A A A A T A G C A C C  
 SEQ. ID. NO. 34 G G G C T G C T G A G G G T G C T A T C A C C A T  
 SEQ. ID. NO. 30 A G T A C T T C C A C G T G G T T G G C G G C A C  
 SEQ. ID. NO. 26 G G G C T G C T G A G G G T G C T A T C A C C A T

SEQ. ID. NO. 38 T G T C T A T C A G C A A G A G G A G A T T G C A  
 SEQ. ID. NO. 34 C G A G C T G G C C T C C T A C C C C A T C A G T  
 SEQ. ID. NO. 30 C A T T G G A T T C G C T C T G A A G G C T G G G  
 SEQ. ID. NO. 26 C G A G C T G G C C T C C T A C C C C A T C A G T

SEQ. ID. NO. 38 G A A G G G G C T G T G A C A A T T T T G C C C A  
 SEQ. ID. NO. 34 G A C T T T G C C T C C T A C T T C C A G A G C C  
 SEQ. ID. NO. 30 C A G A T C C C A G G C T T C C G G G A A T T C C  
 SEQ. ID. NO. 26 G A C T T T G C C T C C T A C T T C C A G A G C C

Figure 9e

SEQ. ID. NO. 38 A A C G A G C A T C A A T T G A T G G A T T T G A  
 SEQ. ID. NO. 34 T G G A C C C T T G G A A C A A C A G C C G G A A  
 SEQ. ID. NO. 30 T G A A G A A G G T C C A T C C C A G G A A G T C  
 SEQ. ID. NO. 26 T G G A C C C T T G G A A C A A C A G C C G G A A

SEQ. ID. NO. 38 T C G A T A C T T T A G A A G C C G A A C T C T T  
 SEQ. ID. NO. 34 C C C C T G G T T C C G T G A A T T C T G G G A G  
 SEQ. ID. NO. 30 T G T C C A C A A T G G T T T T G C C A A G G A G  
 SEQ. ID. NO. 26 C C C C T G G T T C C G T G A A T T C T G G G A G

SEQ. ID. NO. 38 G C C A A T A A T C G A A G A A A T G T G T G G T  
 SEQ. ID. NO. 34 C A G A G G T T C C G C T G C A G C T T C C G G C  
 SEQ. ID. NO. 30 T T T T G G G A A G A A A C A T T T A A C T G C C  
 SEQ. ID. NO. 26 C A G A G G T T C C G C T G C A G C T T C C G G C

SEQ. ID. NO. 38 T T G C A G A A T T C T G G G A G G A G A A T T T  
 SEQ. ID. NO. 34 A G C G A G A C T G C G C A G C C C A C T C T C T  
 SEQ. ID. NO. 30 A C C T C C A A G A A G G T G C A A A A G G A C C  
 SEQ. ID. NO. 26 A G C G A G A C T G C G C A G C C C A C T C T C T

SEQ. ID. NO. 38 T G G C T G C A A G T T A G G A T C A C A T G G G  
 SEQ. ID. NO. 34 C C G G G C T G T G C C C T T T G A G C A G G A G  
 SEQ. ID. NO. 30 T T T A C C T G T G G A C A C C T T T C T G A G A  
 SEQ. ID. NO. 26 C C G G G C T G T G C C C T T T G A A C A G G A G

SEQ. ID. NO. 38 A A A A G G A A C A G T C A T A T A A A G A A A T  
 SEQ. ID. NO. 34 T C C A A G A T C A T G T T T G T G G T C A A T G  
 SEQ. ID. NO. 30 G G T C A C G A A G A A A G T G G C G A C A G G T  
 SEQ. ID. NO. 26 T C C A A G A T C A T G T T T G T G G T C A A T G

SEQ. ID. NO. 38 G C A C A G G G C T G G A G C G A A T T G C T C G  
 SEQ. ID. NO. 34 C A G T G T A C G C C A T G G C C C A T G C G C T  
 SEQ. ID. NO. 30 T T A G C A A C A G C T C G A C A G C C T T C C G  
 SEQ. ID. NO. 26 C A G T G T A C G C C A T G G C C C A T G C G C T

SEQ. ID. NO. 38 G G A T T C A T C T T A T G A A C A G G A A G G A  
 SEQ. ID. NO. 34 C C A C A A C A T G C A C C G T G C C C T C T G C  
 SEQ. ID. NO. 30 A C C C C T C T G T A C A G G G G A T G A G A A C  
 SEQ. ID. NO. 26 C C A C A A C A T G C A C C G T G C C C T C T G C

Figure 9f

SEQ. ID. NO. 38 A A G G T C C A A T T T G T A A T T G A T G C T G  
 SEQ. ID. NO. 34 C C C A A C A C C A C C C G G C T C T G T G A C G  
 SEQ. ID. NO. 30 A T C A G C A G T G T C G A G A C C C C T T A C A  
 SEQ. ID. NO. 26 C C C A A C A C C A C C C G G C T C T G T G A C G

SEQ. ID. NO. 38 T A T A T T C C A T G G C T T A C G C C C T G C A  
 SEQ. ID. NO. 34 C G A T G C G G C C A G T T A A C G G G C G C C G  
 SEQ. ID. NO. 30 T A G A T T A C A C G C A T T T A C G G A T A T C  
 SEQ. ID. NO. 26 C G A T G C G G C C A G T T A A C G G G C G C C G

SEQ. ID. NO. 38 C A A T A T G C A C A A A G A T C T C T G C C C T  
 SEQ. ID. NO. 34 C C T C T A C A A G G A C T T T G T G C T C A A C  
 SEQ. ID. NO. 30 C T A C A A T G T G T A C T T A G C A G T C T A C  
 SEQ. ID. NO. 26 C C T C T A C A A G G A C T T T G T G C T C A A C

SEQ. ID. NO. 38 G G A T A C A T T G G C C T T T G T C C A C G A A  
 SEQ. ID. NO. 34 G T C A A G T T T G A T G C C C C C T T T C G C C  
 SEQ. ID. NO. 30 T C C A T T G C C C A C G C C T T G C A A G A T A  
 SEQ. ID. NO. 26 G T C A A G T T T G A T G C C C C C T T T C G C C

SEQ. ID. NO. 38 T G A G T A C C A T T G A T G G G A A A G A G C T  
 SEQ. ID. NO. 34 C A G C T G A C A C C C A C A A T G A G G T C C G  
 SEQ. ID. NO. 30 T A T A T A C C T G C T T A C C T G G G A G A G G  
 SEQ. ID. NO. 26 C A G C T G A C A C C C A C A A T G A G G T C C G

SEQ. ID. NO. 38 A C T T G G T T A T A T T C G G G C T G T A A A T  
 SEQ. ID. NO. 34 C T T T G A C C G C T T T G G T G A T G G T A T T  
 SEQ. ID. NO. 30 G C T C T T C A C C A A T G G C T C C T G T G C A  
 SEQ. ID. NO. 26 C T T T G A C C G C T T T G G T G A T G G T A T T

SEQ. ID. NO. 38 T T T A A T G G C A G T G C T G G C A C T C C T G  
 SEQ. ID. NO. 34 G G C C G C T A C A A C A T C T T C A C C T A T C  
 SEQ. ID. NO. 30 G A C A T C A A G A A A G T T G A G G C G T G G C  
 SEQ. ID. NO. 26 G G C C G C T A C A A C A T C T T C A C C T A T C

SEQ. ID. NO. 38 T C A C T T T T A A T G A A A A C G G A G A T G C  
 SEQ. ID. NO. 34 T G C G T G C A G G C A G T G G G C G C T A T C G  
 SEQ. ID. NO. 30 A G G T C C T G A A G C A C C T A C G G C A T C T  
 SEQ. ID. NO. 26 T G C G T G C A G G C A G T G G G C G C T A T C G

Figure 9g



SEQ. ID. NO. 38 T C C T G G A C G T T A T G A T A T C T T C C A G  
 SEQ. ID. NO. 34 C T A C C A G A A G G T G G G C T A C T G G G C A  
 SEQ. ID. NO. 30 A A A C T T T A C A A A C A A T A T G G G G G A G  
 SEQ. ID. NO. 26 C T A C C A G A A G G T G G G C T A C T G G G C A

SEQ. ID. NO. 38 T A T C A A A T A A C C A A C A A A A G C A C A G  
 SEQ. ID. NO. 34 G A A G G C T T G A C T C T G G A C A C C A G C C  
 SEQ. ID. NO. 30 C A G G T G A C C T T T G A T G A G T G T G G T G  
 SEQ. ID. NO. 26 G A A G G C T T G A C T C T G G A C A C C A G C C

SEQ. ID. NO. 38 A G T A C A A A G T C A T C G G C C A C T G G A C  
 SEQ. ID. NO. 34 T C A T C C C A T G G G C C T C A C C C T C A G C  
 SEQ. ID. NO. 30 A C C T G G T G G G G A A C T A T T C C A T C A T  
 SEQ. ID. NO. 26 T C A T C C C A T G G G C C T C A C C G T C A G C

SEQ. ID. NO. 38 C A A T C A G C T T C A T C T A A A A G T G G A A  
 SEQ. ID. NO. 34 C G G C C C C C T G C C C G C C T C T C G C T G C  
 SEQ. ID. NO. 30 C A A C T G G C A C C T C T C C C C A G A G G A T  
 SEQ. ID. NO. 26 C G G C C C C C T G G C C G C C T C T C G C T G C

SEQ. ID. NO. 38 G A C A T G C A G T G G G C T C A T A G A G A A C  
 SEQ. ID. NO. 34 A G T G A G C C C T G C C T C C A G A A T G A G G  
 SEQ. ID. NO. 30 G G C T C C A T C G T G T T T A A G G A A G T C G  
 SEQ. ID. NO. 26 A G T G A G C C C T G C C T C C A G A A T G A G G

SEQ. ID. NO. 38 A T A C T C A C C C G G C G T C T G T C T G C A G  
 SEQ. ID. NO. 34 T G A A G A G T G T G C A G C C G G G C G A A G T  
 SEQ. ID. NO. 30 G G T A T T A C A A C G T C T A T G C C A A G A A  
 SEQ. ID. NO. 26 T G A A G A G T G T G C A G C C G G G C G A A G T

SEQ. ID. NO. 38 C C T G C C G T G T A A G C C A G G G G A G A G G  
 SEQ. ID. NO. 34 C T G C T G C T G G C T C T G C A T T C C G T G C  
 SEQ. ID. NO. 30 G G G A G A A A G A C T C T T C A T C A A C G A G  
 SEQ. ID. NO. 26 C T G C T G C T G G C T C T G C A T T C C G T G C

SEQ. ID. NO. 38 A A G A A A A C G G T G A A A G G G G T C C C T T  
 SEQ. ID. NO. 34 C A G C C C T A T G A G T A C C G A T T G G A C G  
 SEQ. ID. NO. 30 G A G A A A A T C C T G T G G A G T G G G T T C T  
 SEQ. ID. NO. 26 C A G C C C T A T G A G T A C C G A T T G G A C G

Figure 9h

SEQ. ID. NO. 38 G C T G C T G G C A C T G T G A A C G C T G T G A  
 SEQ. ID. NO. 34 A A T T C A C T T G C G C T G A T T G T G G C C T  
 SEQ. ID. NO. 30 C C A G G G A G G T G C C C T T C T C C A A C T G  
 SEQ. ID. NO. 26 A A T T C A C T T G C G C T G A T T G T G G C C T

SEQ. ID. NO. 38 A G G T T A C A A C T A C C A G G T G G A T G A G  
 SEQ. ID. NO. 34 G G G C T A C T G G C C C A A T G C C A G C C T G  
 SEQ. ID. NO. 30 C A G C C G A G A C T G C C T G G C A G G G A C C  
 SEQ. ID. NO. 26 G G G C T A C T G G C C C A A T G C C A G C C T G

SEQ. ID. NO. 38 C T G T C C T G T G A A C T T T G C C C T C T G G  
 SEQ. ID. NO. 34 A C T G G C T G C T T C G A A C T G C C C C A G G  
 SEQ. ID. NO. 30 A G G A A A G G G A T C A T T G A G G G G G A G C  
 SEQ. ID. NO. 26 A C T G G C T G C T T C G A A C T G C C C C A G G

SEQ. ID. NO. 38 A T C A G A G A C C C A A C A T G A A C C G C A C  
 SEQ. ID. NO. 34 A G T A C A T C C G C T G G G G C G A T G C C T G  
 SEQ. ID. NO. 30 C C A C C T G C T G C T T T G A G T G T G T G G A  
 SEQ. ID. NO. 26 A G T A C A T C C G C T G G G G C G A T G C C T G

SEQ. ID. NO. 38 A G G C T G C C A G C T T A T C C C C A T C A T C  
 SEQ. ID. NO. 34 G G C T G T G G G A C C T G T C A C C A T C G C C  
 SEQ. ID. NO. 30 G T G T C C T G A T G G G G A G T A T A G T G A T  
 SEQ. ID. NO. 26 G G C T G T G G G A C C T G T C A C C A T C G C C

SEQ. ID. NO. 38 A A A T T G G A G T G G C A T T C T C C C T G G G  
 SEQ. ID. NO. 34 T G C C T C G G T G C C C T G G C C A C C C T C T  
 SEQ. ID. NO. 30 G A G A C A G A T G C C A G T G C C T G T A A C A  
 SEQ. ID. NO. 26 T G C C T C G G T G C C C T G G C C A C C C T G T

SEQ. ID. NO. 38 C T G T G G T G C C T G T G T T T G T T G C A A T  
 SEQ. ID. NO. 34 T T G T G C T G G G T G T C T T T G T G C G G C A  
 SEQ. ID. NO. 30 A G T G C C C A G A T G A C T T C T G G T C C A A  
 SEQ. ID. NO. 26 T T G T G C T G G G T G T C T T T G T G C G G C A

SEQ. ID. NO. 38 A T T G G G A A T C A T C G C C A C C A C C T T T  
 SEQ. ID. NO. 34 C A A T G C C A C A C C A G T G G T C A A G G C C  
 SEQ. ID. NO. 30 T G A G A A C C A C A C C T C C T G C T T C G A A  
 SEQ. ID. NO. 26 C A A T G C C A C A C C A G T G G T C A A G G C C

Figure 9i

SEQ. ID. NO. 38 GTGATCGTGACCTTTGTCCGCTATA  
 SEQ. ID. NO. 34 TCAGGTCTGGGAGCTCTGCTACATCC  
 SEQ. ID. NO. 30 CTGCCCCAGGAGTACATCCGCTGGG  
 SEQ. ID. NO. 26 TCAGGTCTGGGAGCTCTGCTACATCC

SEQ. ID. NO. 38 ATGACACACCTATCTGTGAGGGCTTC  
 SEQ. ID. NO. 34 TGCTGGGTGGTGTCTTCCTCTGCTA  
 SEQ. ID. NO. 30 GCGATGCCCTGGGCTGTGGGACCTGT  
 SEQ. ID. NO. 26 TGCTGGGTGGTGTCTTCCTCTGCTA

SEQ. ID. NO. 38 AGGACGCGAACTTAGTTACGTGCTC  
 SEQ. ID. NO. 34 CTGCATGACCTTCATCTTCATTGCC  
 SEQ. ID. NO. 30 CACCATCGCCTGCTCGGTGCCCTG  
 SEQ. ID. NO. 26 CTGCATGACCTTCATCTTCATTGCC

SEQ. ID. NO. 38 CTAACGGGGATTTTTCTCTGTTATT  
 SEQ. ID. NO. 34 AAGCCATCCACGGCAGTGTGTACCT  
 SEQ. ID. NO. 30 GCCACCCTGTTTGTGCTGGGTGTCT  
 SEQ. ID. NO. 26 AAGCCATCCACGGCAGTGTGTACCT

SEQ. ID. NO. 38 CAATCACGTTTTTTAATGATTGCAGC  
 SEQ. ID. NO. 34 TACGGCGTCTTTGGTTTGGGGCACTGC  
 SEQ. ID. NO. 30 TTGTGCGGCAACAATGCCACACCAGT  
 SEQ. ID. NO. 26 TACGGCGTCTTTGGTTTGGGGCACTGC

SEQ. ID. NO. 38 ACCAGATACAATCATATGCTCCTTC  
 SEQ. ID. NO. 34 CTTCTCTGTCTGCTACTCAGCCCTG  
 SEQ. ID. NO. 30 GGTCAAGGCCTCAGGTCTGGGAGCTC  
 SEQ. ID. NO. 26 CTTCTCTGTCTGCTACTCAGCCCTG

SEQ. ID. NO. 38 CGACGGGTCTTCTCTAGGACTTGGCA  
 SEQ. ID. NO. 34 CTCACCAAGACCAACC GCATTGCAC  
 SEQ. ID. NO. 30 TGCTACATCCTGCTGGGTGGTGTCT  
 SEQ. ID. NO. 26 CTCACCAAGACCAACC GCATTGCAC

SEQ. ID. NO. 38 TGTGTTTTCAGCTATGCAGCCCTTCT  
 SEQ. ID. NO. 34 GCATCTTTCGGTGGGGCCCCGGGAGGG  
 SEQ. ID. NO. 30 TCTCTGTCTACTGCATGACCTTCAT  
 SEQ. ID. NO. 26 GCATCTTTCGGTGGGGCCCCGGGAGGG

Figure 9j

SEQ. ID. NO. 38 G A C C A A A A C A A A C C G T A T C C A C C G A  
 SEQ. ID. NO. 34 T G C C C A G C G G C C A C G C T T C A T C A G T  
 SEQ. ID. NO. 30 C T T C A T T G C C A A G C C A T C C A C G G C A  
 SEQ. ID. NO. 26 T G C C C A G C G G C C A C G C T T C A T C A G T

SEQ. ID. NO. 38 A T A T T T G A G C A G G G G A A G A A A T C T G  
 SEQ. ID. NO. 34 C C T G C C T C A C A G G T G G C C A T C T G C C  
 SEQ. ID. NO. 30 G T G T G T A C C T T A C G G C G T C T T G G T T  
 SEQ. ID. NO. 26 C C T G C C T C A C A G G T G G C C A T C T G C C

SEQ. ID. NO. 38 T C A C A G C G C C C A A G T T C A T T A G T C C  
 SEQ. ID. NO. 34 T G G C A C T T A T C T C G G G C C A G C T G C T  
 SEQ. ID. NO. 30 T G G G C A C T G C C T T C T C T G T C T G C T A  
 SEQ. ID. NO. 26 T G G C A C T T A T C T C G G G C C A G C T G C T

SEQ. ID. NO. 38 A G C A T C T C A G C T G G T G A T C A C C T T C  
 SEQ. ID. NO. 34 C A T C G T G G T C G C C T G G C T G G T G G T G  
 SEQ. ID. NO. 30 C T C A G C C C T G C T C A C C A A G A C C A A C  
 SEQ. ID. NO. 26 C A T C G T G G T C G C C T G G C T G G T G G T G

SEQ. ID. NO. 38 A G C C T C A T C T C C G T C C A G C T C C T T G  
 SEQ. ID. NO. 34 G A G G C A C C G G G C A C A G G C A A G G A G A  
 SEQ. ID. NO. 30 C G C A T T G C A C G C A T C T T C G G T G G G G  
 SEQ. ID. NO. 26 G A G G C A C C G G G C A C A G G C A A G G A G A

SEQ. ID. NO. 38 G A G T G T T T G T C T G G T T T G T T G T G G A  
 SEQ. ID. NO. 34 C A G C C C C C G A A C G G C G G G A G G T G G T  
 SEQ. ID. NO. 30 C C C G G G A G G G T G C C C A G C G G C C A C G  
 SEQ. ID. NO. 26 C A G C C C C C G A A C G G C G G G A G G T G G T

SEQ. ID. NO. 38 T C C C C C C C A C A T C A T C A T T G A C T A T  
 SEQ. ID. NO. 34 G A C A C T G C G C T G C A A C C A C C G C G A T  
 SEQ. ID. NO. 30 C T T C A T C A G T C C T G C C T C A C A G G T G  
 SEQ. ID. NO. 26 G A C A C T G C G C T G C A A C C A C C G C G A T

SEQ. ID. NO. 38 G G A G A G C A G C G G A C A C T A G A T C C A G  
 SEQ. ID. NO. 34 G C A A G T A T G T T G G G C T C G C T G G C C T  
 SEQ. ID. NO. 30 G C C A T C T G C C T G G G C A C T T A T C T C G G  
 SEQ. ID. NO. 26 G C A A G T A T G T T G G G C T C G C T G G C C T

Figure 9k

SEQ. ID. NO. 38 A G A A G G C C A G G G G A G T G C T C A A G T G  
 SEQ. ID. NO. 34 A C A A T G T G C T C C T C A T C G C G C T C T G  
 SEQ. ID. NO. 30 G C C A G C T G C T C A T C G T G G T C G C C T G  
 SEQ. ID. NO. 26 A C A A T G T G C T C C T C A T C G C G C T C T G

SEQ. ID. NO. 38 T G A C A T T T C T G A T C T C T C A C T C A T T  
 SEQ. ID. NO. 34 C A C G C T T T A T G C C T T C A A G A C T C G C  
 SEQ. ID. NO. 30 G C T G G T G G T G G A G G C A C C G G G C A C A  
 SEQ. ID. NO. 26 C A C G C T T T A T G C C T T C A A T A C T C G C

SEQ. ID. NO. 38 T G T T C A C T T G G A T A C A G T A T C C T C T  
 SEQ. ID. NO. 34 A A G T G C C C C G A A A A C T T C A A C G A G G  
 SEQ. ID. NO. 30 G G C A A G G A G A C A G C C C C C G A A C G G C  
 SEQ. ID. NO. 26 A A G T G C C C C G A A A A C T T C A A C G A G G

SEQ. ID. NO. 38 T G A T G G T C A C T T G T A C T G T T T A T G C  
 SEQ. ID. NO. 34 C C A A G T T C A T T G G C T T C A C C A T G T A  
 SEQ. ID. NO. 30 G G G A G G T G G T G A C A C T G C G C T G C A A  
 SEQ. ID. NO. 26 C C A A G T T C A T T G G C T T C A C C A T G T A

SEQ. ID. NO. 38 C A T T A A A A C G A G A G G T G T C C C A G A G  
 SEQ. ID. NO. 34 C A C C A C C T G C A T C A T C T G G C T G G C A  
 SEQ. ID. NO. 30 C C A C C G C G A T G C A A G T A T G T T G G G C  
 SEQ. ID. NO. 26 C A C C A C C T G C A T C A T C T G G C T G G C A

SEQ. ID. NO. 38 A C T T T C A A T G A A G C C A A A C C T A T T G  
 SEQ. ID. NO. 34 T T C C T G C C C A T C T T C T A T G T C A C C T  
 SEQ. ID. NO. 30 T C G C T G G C C T A C A A T G T G C T C C T C A  
 SEQ. ID. NO. 26 T T G T T G C C C A T C T T C T A T G T C A C C T

SEQ. ID. NO. 38 G A T T T A C C A T G T A T A C C A C C T G C A T  
 SEQ. ID. NO. 34 C C A G T G A C T A C C G G G T A C A G A C C A C  
 SEQ. ID. NO. 30 T C G C G C T C T G C A C G C T T T A T G C C T T  
 SEQ. ID. NO. 26 C C A G T G A C T A C C G G G T A C A G A C C A C

SEQ. ID. NO. 38 C A T T T G G T T A G C T T T C A T C C C C A T C  
 SEQ. ID. NO. 34 C A C C A T G T G C G T G T C A G T C A G C C T C  
 SEQ. ID. NO. 30 C A A T A C T C G C A A G T G C C C C G A A A A C  
 SEQ. ID. NO. 26 C A C C A T G T G C G T G T C A G T C A G C C T C

Figure 91

SEQ. ID. NO. 38 T T T T T T G G T A C A G C C C A G T C A G C A G  
 SEQ. ID. NO. 34 A G C G G C T C C G T G G T G C T T G G C T G C C  
 SEQ. ID. NO. 30 T T C A A C G A G G C C A A G T T C A T T G G C T  
 SEQ. ID. NO. 26 A G C G G C T C C G T G G T G C T T G G C T G C C

SEQ. ID. NO. 38 A A A A G A T G T A C A T C C A G A C A A C A A C  
 SEQ. ID. NO. 34 T C T T T G C G C C C A A G C T G C A C A T C A T  
 SEQ. ID. NO. 30 T C A C C A T G T A C A C C A C C T G C A T C A T  
 SEQ. ID. NO. 26 T C T T T G C G C C C A A G C T G C A C A T C A T

SEQ. ID. NO. 38 A C T T A C T G T C T C C A T G A G T T T A A G T  
 SEQ. ID. NO. 34 C C T C T T C C A G C C G C A G A A G A A C A C C  
 SEQ. ID. NO. 30 C T G G C T G G C A T T G T T G C C C A T C T T C  
 SEQ. ID. NO. 26 C C T C T T C C A G C C G C A G A A G A A C G T G

SEQ. ID. NO. 38 G C T T C A G T A T C T C T G G G C A T G C T C T  
 SEQ. ID. NO. 34 A T C G A G G A G G T G C G T T G C A G C A C C G  
 SEQ. ID. NO. 30 T A T G T C A C C T C C A G T G A C T A C C G G G  
 SEQ. ID. NO. 26 G T T A G C C A C C G G G C A C C C A C C A G C C

SEQ. ID. NO. 38 A T A T G C C C A A G G T T T A T A T T A T A A T  
 SEQ. ID. NO. 34 C A G C T C A C G C T T T C A A G G T G G C T G C  
 SEQ. ID. NO. 30 T A C A G A C C A C C A C C A T G T G C G T G T C  
 SEQ. ID. NO. 26 G C T T T G G C A G T G C T G C T G C C A G G G C

SEQ. ID. NO. 38 T T T T C A T C C A G A A C A G A A T A C C A T C  
 SEQ. ID. NO. 34 C C G G G C C A C G C T G C G C C G C A G C A A C  
 SEQ. ID. NO. 30 A G T C A G C C T C A G C G G C T C C G T G G T G  
 SEQ. ID. NO. 26 C A G C T C C A G C C T T G G C C A A G G G T C T

SEQ. ID. NO. 38 G A G G A G G T G C G T T G C A G C A C C G C A G  
 SEQ. ID. NO. 34 G T C T C C C G C A A G C G G T C C A G C A G C C  
 SEQ. ID. NO. 30 C T T G G C T G C C T C T T T G C G C C C A A G C  
 SEQ. ID. NO. 26 G G C T C C C A G T T T G T C C C C A C T G T T T

SEQ. ID. NO. 38 C T C A C G C T T T C A A G G T G G C T G C C C G  
 SEQ. ID. NO. 34 T T G G A G G C T C C A C G G G A T C C A C C C C  
 SEQ. ID. NO. 30 T G C A C A T C A T C C T C T T C C A G C C G C A  
 SEQ. ID. NO. 26 G C A A T G G C C G T G A G G T G G T G G A C T C

Figure 9m

SEQ. ID. NO. 38 G G C C A C G C T G C G C C G C A G C A A C G T C  
 SEQ. ID. NO. 34 C T C C T C C T C C A T C A G C A G C A A G A G C  
 SEQ. ID. NO. 30 G A A G A A C G T G G T T A G C C A C C G G G C A  
 SEQ. ID. NO. 26 G A C A A C G T C A T C G C T T

SEQ. ID. NO. 38 T C C C G C A A A G C G G T C C A G C A G C C T T G  
 SEQ. ID. NO. 34 A A C A G C G A A G A C C C A T T C C C A C A G C  
 SEQ. ID. NO. 30 C C C A C C A G C C G C T T T G G C A G T G C T G  
 SEQ. ID. NO. 26

SEQ. ID. NO. 38 G A G G C T C C A C G G G A T C C A C C C C C T C  
 SEQ. ID. NO. 34 C C G A G A G G C A G A A G C A G C A G C A G C C  
 SEQ. ID. NO. 30 C T G C C A G G G C C A G C T C C A G C C T T G G  
 SEQ. ID. NO. 26

SEQ. ID. NO. 38 C T C C T C C A T C A G C A G C A A G A G C A A C  
 SEQ. ID. NO. 34 G C T G G C C C T A A C C C A G C A A G A G C A G  
 SEQ. ID. NO. 30 C C A A G G G T C T G G C T C C C A G T T T G T C  
 SEQ. ID. NO. 26

SEQ. ID. NO. 38 A G C G A A G A C C C A T T C C C A C A G C C C G  
 SEQ. ID. NO. 34 C A G C A G C A G C C C C T G A C C C T C C C A C  
 SEQ. ID. NO. 30 C C C A C T G T T T G C A A T G G C C G T G A G G  
 SEQ. ID. NO. 26

SEQ. ID. NO. 38 A G A G G C A G A A G C A G C A G C A G C C G C T  
 SEQ. ID. NO. 34 A G C A G C A A C G A T C T C A G C A G C A G C C  
 SEQ. ID. NO. 30 T G G T G G A C T C G A C A A C G T C A T C G C T  
 SEQ. ID. NO. 26

SEQ. ID. NO. 38 G G C C C T A A C C C A G C A A G A G C A G C A G  
 SEQ. ID. NO. 34 C A G A T G C A A G C A G A A G G T C A T C T T T  
 SEQ. ID. NO. 30 T  
 SEQ. ID. NO. 26

SEQ. ID. NO. 38 C A G C A G C C C C T G A C C C T C C C A C A G C  
 SEQ. ID. NO. 34 G G C A G C G G C A C G G T C A C C T T C T C A C  
 SEQ. ID. NO. 30  
 SEQ. ID. NO. 26

Figure 9n

SEQ. ID. NO. 38 A G C A A C G A T C T C A G C A G C A G C C C A G  
SEQ. ID. NO. 34 T G A G C T T T G A T G A G C C T C A G A A G A A  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 A T G C A A G C A G A A G G T C A T C T T T G G C  
SEQ. ID. NO. 34 C G C C A T G G C C C A C G G G A A T T C T A C G  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 A G C G G C A C G G T C A C C T T C T C A C T G A  
SEQ. ID. NO. 34 C A C C A G A A C T C C C T G G A G G C C C A G A  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 G C T T T G A T G A G C C T C A G A A G A A C G C  
SEQ. ID. NO. 34 A A A G C A G C G A T A C G C T G A C C C G A C A  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 C A T G G C C C A C G G G A A T T C T A C G C A C  
SEQ. ID. NO. 34 C C A G C C A T T A C T C C C G C T G C A G T G C  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 C A G A A C T C C C T G G A G G C C C A G A A A A  
SEQ. ID. NO. 34 G G G G A A A C G G A C T T A G A T C T G A C C G  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 G C A G C G A T A C G C T G A C C C G A C A C C A  
SEQ. ID. NO. 34 T C C A G G A A A C A G G T C T G C A A G G A C C  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 G C C A T T A C T C C C G C T G C A G T G C G G G  
SEQ. ID. NO. 34 T G T G G G T G G A G A C C A G C G G C C A G A G  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

Figure 9o



SEQ. ID. NO. 38 G A A A C G G A C T T A G A T C T G A C C G T C C  
SEQ. ID. NO. 34 G T G G A G G A C C C T G A A G A G T T G T C C C  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 A G G A A A C A G G T C T G C A A G G A C C T G T  
SEQ. ID. NO. 34 C A G C A C T T G T A G T G T C C A G T T C A C A  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 G G G T G G A G A C C A G C G G C C A G A G G T G  
SEQ. ID. NO. 34 G A G C T T T G T C A T C A G T G G T G G A G G C  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 G A G G A C C C T G A A G A G T T G T C C C C A G  
SEQ. ID. NO. 34 A G C A C T G T T A C A G A A A C G T A G T G A  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 C A C T T G T A G T G T C C A G T T C A C A G A G  
SEQ. ID. NO. 34 A T T C A  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 C T T T G T C A T C A G T G G T G G A G G C A G C  
SEQ. ID. NO. 34  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 A C T G T T A C A G A A A C G T A G T G A A T T  
SEQ. ID. NO. 34  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

SEQ. ID. NO. 38 C A  
SEQ. ID. NO. 34  
SEQ. ID. NO. 30  
SEQ. ID. NO. 26

Figure 9p

# ClustalW Formatted Alignments

SEQ. ID. NO. 39 M V C E G K R S A S C P C F F L L T A K F Y W I L  
 SEQ. ID. NO. 35 M G S L L A L P A L L L L W G A V A E G P A K K V  
 SEQ. ID. NO. 31 M A F Y S C C W V L L A L T W H T S A Y G P D Q R  
 SEQ. ID. NO. 27 M G S L L A L L A L L P L W G A V A E G P A K K V

SEQ. ID. NO. 39 T M M Q R T H S Q E Y A H S I R V D G D I I L G G  
 SEQ. ID. NO. 35 L T L E G D L V L G G L F P V H Q K G G P A E D C  
 SEQ. ID. NO. 31 A Q K K G D I I L G G L F P I H F G V A A K D Q D  
 SEQ. ID. NO. 27 L T L E G D L V L G G L F P V H Q K G G P A E D C

SEQ. ID. NO. 39 L F P V H A K G E R G V P C G E L K K E K G I H R  
 SEQ. ID. NO. 35 G P V N E H R G I Q R L E A M L F A L D R I N R D  
 SEQ. ID. NO. 31 L K S R P E S V E C I R Y N F R G F R W L Q A M I  
 SEQ. ID. NO. 27 G P V N E H R G I Q R L E A M L F A L D R I N R D

SEQ. ID. NO. 39 L E A M L Y A I D Q I N K D P D L L S N I T L G V  
 SEQ. ID. NO. 35 P H L L P G V R L G A H I L D S C S K D T H A L E  
 SEQ. ID. NO. 31 F A I E E I N S S P A L L P N L T L G Y R I F D T  
 SEQ. ID. NO. 27 P H L L P G V R L G A H I L D S C S K D T H A L E

SEQ. ID. NO. 39 R I L D T C S R D T Y A L E Q S L T F V Q A L I E  
 SEQ. ID. NO. 35 Q A L D F V R A S L S R G A D G S R H I C P D G S  
 SEQ. ID. NO. 31 C N T V S K A L E A T L S F V A Q N K I D S L N L  
 SEQ. ID. NO. 27 Q A L D F V R A S L S R G A D G S R H I C P D G S

SEQ. ID. NO. 39 K D A S D V K C A N G D P P I F T K P D K I S G V  
 SEQ. ID. NO. 35 Y A T H G D A P T A I T G V I G G S Y S D V S I Q  
 SEQ. ID. NO. 31 D E F C N C S E H I P S T I A V V G A T G S G V S  
 SEQ. ID. NO. 27 Y A T H G D A P T A I T G V I G G S Y S D V S I Q

SEQ. ID. NO. 39 I G A A A S S V S I M V A N I L R L F K I P Q I S  
 SEQ. ID. NO. 35 V A N L L R L F Q I P Q I S Y A S T S A K L S D K  
 SEQ. ID. NO. 31 T A V A N L L G L F Y I P Q V S Y A S S S R L L S  
 SEQ. ID. NO. 27 V A N L L R L F Q I P Q I S Y A S T S A K L S D K

SEQ. ID. NO. 39 Y A S T A P E L S D N T R Y D F F S R V V P P D S  
 SEQ. ID. NO. 35 S R Y D Y F A R T V P P D F F Q A K A M A E I L R  
 SEQ. ID. NO. 31 N K N Q F K S F L R T I P N D E H Q A T A M A D I  
 SEQ. ID. NO. 27 S R Y D Y F A R T V P P D F F Q A K A M A E I L R

Figure 10a

SEQ. ID. NO. 39 YQAQAMVDIVTALGWNYVSTLASEG  
 SEQ. ID. NO. 35 FFNWTYVSTVASSEGDYGETGIEAFE  
 SEQ. ID. NO. 31 IEYFRWNWVGTTAAADDDYGRPGIEK  
 SEQ. ID. NO. 27 FFNWTYVSTEASEGDYGETGIEAFE

SEQ. ID. NO. 39 NYGESGVEAFTQISREIGGVCI AQ S  
 SEQ. ID. NO. 35 LEARARNICVATSEKVGGRAMSRAAF  
 SEQ. ID. NO. 31 FREEAEERDIDIDFSELISQYSDEE  
 SEQ. ID. NO. 27 LEARARNICVATSEKVGGRAMSRAAF

SEQ. ID. NO. 39 QKIPREPRPGEFEKI IKRLLETPNA  
 SEQ. ID. NO. 35 EGVVRALLQKPSARVAVLFTRSEDA  
 SEQ. ID. NO. 31 EIQHVVEVIQNSTAKVIVVFSSGPD  
 SEQ. ID. NO. 27 EGVVRALLQKPSARVAVLFTRSEDA

SEQ. ID. NO. 39 RAVIMFANEDDIRRILEAAKKLNQS  
 SEQ. ID. NO. 35 REL LAASQRLNASFTWVASD GWGAL  
 SEQ. ID. NO. 31 LEPLIKEIVRRNITGKIWLASEAWA  
 SEQ. ID. NO. 27 REL LAASQRLNASFTWVASD GWGAL

SEQ. ID. NO. 39 GHFLWIGSDSWGSKIAPVYQQEEIA  
 SEQ. ID. NO. 35 ESVVAGSEGA AEGAITIELASYPIS  
 SEQ. ID. NO. 31 SSSLIAMPQYFHVVG GTIGFALKAG  
 SEQ. ID. NO. 27 ESVVAGSEGA AEGAITIELASYPIS

SEQ. ID. NO. 39 EGAVTILPKRASIDGFDRYFRSRTL  
 SEQ. ID. NO. 35 DFASYFQSLDPWNNNSRNPWFREFWE  
 SEQ. ID. NO. 31 QIPGFREFLKKVHPRKSVHNGFAKE  
 SEQ. ID. NO. 27 DFASYFQSLDPWNNNSRNPWFREFWE

SEQ. ID. NO. 39 ANNRRNVWF AEFW EENFGCKLGSHG  
 SEQ. ID. NO. 35 QRFRC SFRQRDCAAHSLRAVPFEQE  
 SEQ. ID. NO. 31 FWEE TFNCHLQEGAKGPLPVD TFLR  
 SEQ. ID. NO. 27 QRFRC SFRQRDCAAHSLRAVPFEQE

SEQ. ID. NO. 39 KRN SHIKKCTGLERIARDSSSYEQEG  
 SEQ. ID. NO. 35 SKIMFVVNAVYAMAHALHNMHRA LC  
 SEQ. ID. NO. 31 GHEESGDRFSNSSTA FRPLCTGDEN  
 SEQ. ID. NO. 27 SKIMFVVNAVYAMAHALHNMHRA LC

Figure 10b

SEQ. ID. NO. 39 K V Q F V I D A V Y S M A Y A L H N M H K D L C P  
 SEQ. ID. NO. 35 P N T T R L C D A M R P V N G R R L Y K D F V L N  
 SEQ. ID. NO. 31 I S S V B T P Y I D Y T H L R I S Y N V Y L A V Y  
 SEQ. ID. NO. 27 P N T T R L C D A M R P V N G R R L Y K D F V L N

SEQ. ID. NO. 39 G Y I G L C P R M S T I D G K E L L G Y I R A V N  
 SEQ. ID. NO. 35 V K F D A P F R P A D T H N E V R F D R F G D G I  
 SEQ. ID. NO. 31 S I A H A L Q D I Y T C L P G R G L F T N G S C A  
 SEQ. ID. NO. 27 V K F D A P F R P A D T H N E V R F D R F G D G I

SEQ. ID. NO. 39 F N G S A G T P V T F N E N G D A P G R Y D I F Q  
 SEQ. ID. NO. 35 G R Y N I F T Y L R A G S G R Y R Y Q K V G Y W A  
 SEQ. ID. NO. 31 D I K K V E A W Q V L K H L R H L N F T N N M G E  
 SEQ. ID. NO. 27 G R Y N I F T Y L R A G S G R Y R Y Q K V G Y W A

SEQ. ID. NO. 39 Y Q I T N K S T E Y K V I G H W T N Q L H L K V E  
 SEQ. ID. NO. 35 E G L T L D T S L I P W A S P S A G P L P A S R C  
 SEQ. ID. NO. 31 Q V T F D E C G D L V G N Y S I I N W H L S P E D  
 SEQ. ID. NO. 27 E G L T L D T S L I P W A S P S A G P L A A S R C

SEQ. ID. NO. 39 D M Q W A H R E H T H P A S V C S L P C K P G E R  
 SEQ. ID. NO. 35 S E P C L Q N E V K S V Q P G E V C C W L C I P C  
 SEQ. ID. NO. 31 G S I V F K E V G Y Y N V Y A K K G E R L F I N E  
 SEQ. ID. NO. 27 S E P C L Q N E V K S V Q P G E V C C W L C I P C

SEQ. ID. NO. 39 K K T V K G V P C C W H C E R C E G Y N Y Q V D E  
 SEQ. ID. NO. 35 Q P Y E Y R L D E F T C A D C G L G Y W P N A S L  
 SEQ. ID. NO. 31 E K I L W S G F S R E V P F S N C S R D C L A G T  
 SEQ. ID. NO. 27 Q P Y E Y R L D E F T C A D C G L G Y W P N A S L

SEQ. ID. NO. 39 L S C E L C P L D Q R P N M N R T G C Q L I P I I  
 SEQ. ID. NO. 35 T G C F E L P Q E Y I R W G D A W A V G P V T I A  
 SEQ. ID. NO. 31 R K G I I E G E P T C C F E C V E C P D G E Y S D  
 SEQ. ID. NO. 27 T G C F E L P Q E Y I R W G D A W A V G P V T I A

SEQ. ID. NO. 39 K L E W H S P W A V V P V F V A I L G I I A T T F  
 SEQ. ID. NO. 35 C L G A L A T L F V L G V F V R H N A T P V V K A  
 SEQ. ID. NO. 31 B T D A S A C N K C P D D F W S N E N H T S C F E  
 SEQ. ID. NO. 27 C L G A L A T L F V L G V F V R H N A T P V V K A

Figure 10c

SEQ. ID. NO. 39 V I V T F V R Y N D T P I V R A S G R E L S Y V L  
 SEQ. ID. NO. 35 S G R E L C Y I L L G G V F L C Y C M T F I F I A  
 SEQ. ID. NO. 31 L P Q E Y I R W G D A W A V G P V T I A C L G A L  
 SEQ. ID. NO. 27 S G R E L C Y I L L G G V F L C Y C M T F I F I A

SEQ. ID. NO. 39 L T G I F L C Y S I T F L M I A A P D T I I C S F  
 SEQ. ID. NO. 35 K P S T A V C T L R R L G L G T A F S V C Y S A L  
 SEQ. ID. NO. 31 A T L F V L G V F V R H N A T P V V K A S G R E L  
 SEQ. ID. NO. 27 K P S T A V C T L R R L G L G T A F S V C Y S A L

SEQ. ID. NO. 39 R R V F L G L G M C F S Y A A L L T K T N R I H R  
 SEQ. ID. NO. 35 L T K T N R I A R I F G G A R E G A Q R P R F I S  
 SEQ. ID. NO. 31 C Y I L L G G V F L C Y C M T F I F I A K P S T A  
 SEQ. ID. NO. 27 L T K T N R I A R I F G G A R E G A Q R P R F I S

SEQ. ID. NO. 39 I F E Q G K K S V T A P K F I S P A S Q L V I T F  
 SEQ. ID. NO. 35 P A S Q V A I C L A L I S G Q L L I V V A W L V V  
 SEQ. ID. NO. 31 V C T L R R L G L G T A F S V C Y S A L L T K T N  
 SEQ. ID. NO. 27 P A S Q V A I C L A L I S G Q L L I V V A W L V V

SEQ. ID. NO. 39 S L I S V Q L L G V F V W F V V D P P H I I I D Y  
 SEQ. ID. NO. 35 E A P G T G K E T A P E R R E V V T L R C N H R D  
 SEQ. ID. NO. 31 R I A R I F G G A R E G A Q R P R F I S P A S Q V  
 SEQ. ID. NO. 27 E A P G T G K E T A P E R R E V V T L R C N H R D

SEQ. ID. NO. 39 G E Q R T L D P E K A R G V L K C D I S D L S L I  
 SEQ. ID. NO. 35 A S M L G S L A Y N V L L I A L C T L Y A F K T R  
 SEQ. ID. NO. 31 A I C L A L I S G Q L L I V V A W L V V E A P G T  
 SEQ. ID. NO. 27 A S M L G S L A Y N V L L I A L C T L Y A F N T R

SEQ. ID. NO. 39 C S L G Y S I L L M V T C T V Y A I K T R G V P E  
 SEQ. ID. NO. 35 K C P E N F N E A K F I G F T M Y T T C I I W L A  
 SEQ. ID. NO. 31 G K E T A P E R R E V V T L R C N H R D A S M L G  
 SEQ. ID. NO. 27 K C P E N F N E A K F I G F T M Y T T C I I W L A

SEQ. ID. NO. 39 T F N E A K P I G F T M Y T T C I I W L A F I P I  
 SEQ. ID. NO. 35 F L P I F Y V T S S D Y R V Q T T T M C V S V S L  
 SEQ. ID. NO. 31 S L A Y N V L L I A L C T L Y A F N T R K C P E N  
 SEQ. ID. NO. 27 L L P I F Y V T S S D Y R V Q T T T M C V S V S L

Figure 10d

SEQ. ID. NO. 39 P F G T A Q S A E K M Y I Q T T T L T V S M S L S  
 SEQ. ID. NO. 35 S G S V V L G C L F A P K L H I I L F Q P Q K N T  
 SEQ. ID. NO. 31 F N E A K F I G F T M Y T T C I I W L A L L P I F  
 SEQ. ID. NO. 27 S G S V V L G C L F A P K L H I I L F Q P Q K N

SEQ. ID. NO. 39 A S V S L G M L Y M P K V Y I I I F H P E Q N T I  
 SEQ. ID. NO. 35 I E E V R C S T A A H A F K V A A R A T L R R S N  
 SEQ. ID. NO. 31 Y V T S S D Y R V Q T T T M C V S V S L S G S V V  
 SEQ. ID. NO. 27

SEQ. ID. NO. 39 E E V R C S T A A H A F K V A A R A T L R R S N V  
 SEQ. ID. NO. 35 V S R K R S S S L G G S T G S T P S S S I S S K S  
 SEQ. ID. NO. 31 L G C L F A P K L H I I L F Q P Q K N V V S H R A  
 SEQ. ID. NO. 27

SEQ. ID. NO. 39 S R K R S S S L G G S T G S T P S S S I S S K S N  
 SEQ. ID. NO. 35 N S E D P F P Q P E R Q K Q Q Q P L A L T Q Q E Q  
 SEQ. ID. NO. 31 P T S R F G S A A R A S S S L G Q G S G S Q F V  
 SEQ. ID. NO. 27

SEQ. ID. NO. 39 S E D P F P Q P E R Q K Q Q Q P L A L T Q Q E Q Q  
 SEQ. ID. NO. 35 Q Q Q P L T L P Q Q Q R S Q Q Q P R C K Q K V I F  
 SEQ. ID. NO. 31 P T V C N G R E V V D S T T S S L  
 SEQ. ID. NO. 27

SEQ. ID. NO. 39 Q Q P L T L P Q Q Q R S Q Q Q P R C K Q K V I F G  
 SEQ. ID. NO. 35 G S G T V T F S L S F D E P Q K N A M A H G N S T  
 SEQ. ID. NO. 31  
 SEQ. ID. NO. 27

SEQ. ID. NO. 39 S G T V T F S L S F D E P Q K N A M A H G N S T H  
 SEQ. ID. NO. 35 H Q N S L E A Q K S S D T L T R H Q P L L P L Q C  
 SEQ. ID. NO. 31  
 SEQ. ID. NO. 27

SEQ. ID. NO. 39 Q N S L E A Q K S S D T L T R H Q P L L P L Q C G  
 SEQ. ID. NO. 35 G E T D L D L T V Q E T G L Q G P V G G D Q R P E  
 SEQ. ID. NO. 31  
 SEQ. ID. NO. 27

Figure 10e

SEQ. ID. NO. 39 E T D L D L T V Q E T G L Q G P V G G D Q R P E V  
SEQ. ID. NO. 35 V E D P E E L S P A L V V S S S Q S F V I S G G G  
SEQ. ID. NO. 31  
SEQ. ID. NO. 27

SEQ. ID. NO. 39 E D P E E L S P A L V V S S S Q S F V I S G G G S  
SEQ. ID. NO. 35 S T V T E N V V N S  
SEQ. ID. NO. 31  
SEQ. ID. NO. 27

SEQ. ID. NO. 39 T V T E N V V N S  
SEQ. ID. NO. 35  
SEQ. ID. NO. 31  
SEQ. ID. NO. 27

# ClustalW Formatted Alignments

SEQ. ID. NO. 40 A T G G T A T G C G A G G G A A A G C G A T C A G  
 SEQ. ID. NO. 46 A T G G G A T C G C T G C T T G C G C T C C C G G  
 SEQ. ID. NO. 36 A T G G G A T C G C T G C T T G C G C T C C C G G  
 SEQ. ID. NO. 32 A T G G C A T T T T A T A G C T G C T G C T G G G

SEQ. ID. NO. 40 C C T C T T G C C C T T G T T T C T T C C T C T T  
 SEQ. ID. NO. 46 C A C T G C T G C T G C T G T G G G G T G C T G T  
 SEQ. ID. NO. 36 C A C T G C T G C T G C T G T G G G G T G C T G T  
 SEQ. ID. NO. 32 T C C T C T T G G C A C T C A C C T G G C A C A C

SEQ. ID. NO. 40 G A C C G C C A A G T T C T A C T G G A T C C T C  
 SEQ. ID. NO. 46 G G C T G A G G G C C C A G C C A A G A A A G G T G  
 SEQ. ID. NO. 36 G G C T G A G G G C C C A G C C A A G A A A G G T G  
 SEQ. ID. NO. 32 C T C T G C C T A C G G G C C A G A C C A G C G A

SEQ. ID. NO. 40 A C A A T G A T G C A A A G A A A C T C A C A G C C  
 SEQ. ID. NO. 46 C T G A C C C T G G A G G G A G A C T T G G T G C  
 SEQ. ID. NO. 36 C T G A C C C T G G A G G G A G A C T T G G T G C  
 SEQ. ID. NO. 32 G C C C A A A A G A A A G G G G G A C A T T A T C C

SEQ. ID. NO. 40 A G G A G T A T G C C C A T T C C A T A C G G G T  
 SEQ. ID. NO. 46 T G G G T G G G C T G T T C C C A G T G C A C C A  
 SEQ. ID. NO. 36 T G G G T G G G C T G T T C C C A G T G C A C C A  
 SEQ. ID. NO. 32 T T G G G G G G C T C T T T C C T A T T C A T T T

SEQ. ID. NO. 40 G G A T G G G G A C A T T A T T T T G G G G G G T  
 SEQ. ID. NO. 46 G A A G G G C G G C C C A G C A G A G G A C T G T  
 SEQ. ID. NO. 36 G A A G G G C G G C C C A G C A G A G G A C T G T  
 SEQ. ID. NO. 32 T G G A G T A G C A G C T A A A G A T C A A G A T

SEQ. ID. NO. 40 C T C T T C C C T G T C C A C G C A A A G G G A G  
 SEQ. ID. NO. 46 G G T C C T G T C A A T G A G C A C C G T G G C A  
 SEQ. ID. NO. 36 G G T C C T G T C A A T G A G C A C C G T G G C A  
 SEQ. ID. NO. 32 C T C A A A T C A A G G C C G G A G T C T G T G G

SEQ. ID. NO. 40 A G A G A G G G G T G C C T T G T G G G G A G C T  
 SEQ. ID. NO. 46 T C C A G C G C C T G G A G G C C A T G C T T T T  
 SEQ. ID. NO. 36 T C C A G C G C C T G G A G G C C A T G C T T T T  
 SEQ. ID. NO. 32 A A T G T A T C A G G T A T A A T T T C C G T G G

Figure 11a



SEQ. ID. NO. 40 G A A G A A G G A A A A G G G G A T T C A C A G A  
 SEQ. ID. NO. 46 T G C A C T G G A C C G C A T C A A C C G T G A C  
 SEQ. ID. NO. 36 T G C A C T G G A C C G C A T C A A C C G T G A C  
 SEQ. ID. NO. 32 G T T T C G C T G G T T A C A G G C T A T G A T A

SEQ. ID. NO. 40 C T G G A G G C C A T G C T T T A T G C A A T T G  
 SEQ. ID. NO. 46 C C G C A C C T G C T G C C T G G C G T G C G C C  
 SEQ. ID. NO. 36 C C G C A C C T G C T G C C T G G C G T G C G C C  
 SEQ. ID. NO. 32 T T T G C C A T A G A G G A G A T A A A C A G C A

SEQ. ID. NO. 40 A C C A G A T T A A C A A A G G A C C C T G A T C T  
 SEQ. ID. NO. 46 T G G G T G C A C A C A T C C T C G A C A G T T G  
 SEQ. ID. NO. 36 T G G G T G C A C A C A T C C T C G A C A G T T G  
 SEQ. ID. NO. 32 G C C C A G C C C T T C T T C C C A A C T T G A C

SEQ. ID. NO. 40 C C T T T C C A A C A T C A C T C T G G G T G T C  
 SEQ. ID. NO. 46 C T C C A A G G A C A C A C A T G C G C T G G A G  
 SEQ. ID. NO. 36 C T C C A A G G A C A C A C A T G C G C T G G A G  
 SEQ. ID. NO. 32 G C T G G G A T A C A G G A T A T T T G A C A C T

SEQ. ID. NO. 40 C G C A T C C T C G A C A C G T G C T C T A G G G  
 SEQ. ID. NO. 46 C A G G C A C T G G A C T T T T G T G C G T G C C T  
 SEQ. ID. NO. 36 C A G G C A C T G G A C T T T T G T G C G T G C C T  
 SEQ. ID. NO. 32 T G C A A C A C C G T T T C T A A G G C C T T G G

SEQ. ID. NO. 40 A C A C C T A T G C T T T T G G A G C A G T C T C T  
 SEQ. ID. NO. 46 C A C T C A G C C G T G G T G C T G A T G G C T C  
 SEQ. ID. NO. 36 C A C T C A G C C G T G G T G C T G A T G G C T C  
 SEQ. ID. NO. 32 A A G C C A C C C T G A G T T T T G T T G C T C A

SEQ. ID. NO. 40 A A C A T T C G T G C A G G C A T T A A T A G A G  
 SEQ. ID. NO. 46 A C G C C A C A T C T G C C C C G A C G G C T C T  
 SEQ. ID. NO. 36 A C G C C A C A T C T G C C C C G A C G G C T C T  
 SEQ. ID. NO. 32 A A A C A A A A T T G A T T C T T T G A A C C T T

SEQ. ID. NO. 40 A A A G A T G C T T C G G A T G T G A A G T G T G  
 SEQ. ID. NO. 46 T A T G C G A C C C A T G G T G A T G C T C C C A  
 SEQ. ID. NO. 36 T A T G C G A C C C A T G G T G A T G C T C C C A  
 SEQ. ID. NO. 32 G A T G A G T T C T G C A A C T G C T C A G A G C

Figure 11b

SEQ. ID. NO. 40 C T A A T G G A G A T C C A C C C A T T T T C A C  
 SEQ. ID. NO. 46 C T G C C A T C A C T G G T G T T A T T G G C G G  
 SEQ. ID. NO. 36 C T G C C A T C A C T G G T G T T A T T G G C G G  
 SEQ. ID. NO. 32 A C A T T C C C T C T A C G A T T G C T G T G G T

SEQ. ID. NO. 40 C A A G C C C G A C A A G A T T T C T G G C G T C  
 SEQ. ID. NO. 46 T T C C T A C A G T G A T G T C T C C A T C C A G  
 SEQ. ID. NO. 36 T T C C T A C A G T G A T G T C T C C A T C C A G  
 SEQ. ID. NO. 32 G G G A G C A A C T G G C T C A G G C G T C T C C

SEQ. ID. NO. 40 A T A G G T G C T G C A G C A A G C T C C G T G T  
 SEQ. ID. NO. 46 G T G G C C A A C C T C T T G A G G C T A T T T C  
 SEQ. ID. NO. 36 G T G G C C A A C C T C T T G A G G C T A T T T C  
 SEQ. ID. NO. 32 A C G G C A G T G G C A A A T C T G C T G G G G C

SEQ. ID. NO. 40 C C A T C A T G G T T G C T A A C A T T T T A A G  
 SEQ. ID. NO. 46 A G A T C C C A C A G A T T A G C T A C G C C T C  
 SEQ. ID. NO. 36 A G A T C C C A C A G A T T A G C T A C G C C T C  
 SEQ. ID. NO. 32 T C T T C T A C A T T C C C A G G T C A G T T A

SEQ. ID. NO. 40 A C T T T T T A A G A T A C C T C A A A T C A G C  
 SEQ. ID. NO. 46 T A C C A G T G C C A A G C T G A G T G A C A A G  
 SEQ. ID. NO. 36 T A C C A G T G C C A A G C T G A G T G A C A A G  
 SEQ. ID. NO. 32 T G C C T C C T C C A G C A G A C T C C T C A G C

SEQ. ID. NO. 40 T A T G C A T C C A C A G C C C C A G A G C T A A  
 SEQ. ID. NO. 46 T C C C G C T A T G A C T A C T T T G C C C G C A  
 SEQ. ID. NO. 36 T C C C G C T A T G A C T A C T T T G C C C G C A  
 SEQ. ID. NO. 32 A A C A A G A A T C A A T T C A A G T C T T T C C

SEQ. ID. NO. 40 G T G A T A A C A C C A G G T A T G A C T T T T T  
 SEQ. ID. NO. 46 C A G T G C C T C C T G A C T T C T T C C A A G C  
 SEQ. ID. NO. 36 C A G T G C C T C C T G A C T T C T T C C A A G C  
 SEQ. ID. NO. 32 T C C G A A C C A T C C C C A A T G A T G A G C A

SEQ. ID. NO. 40 C T C T C G A G T G G T T C C G C C T G A C T C C  
 SEQ. ID. NO. 46 C A A G G C C A T G G C T G A G A T T C T C C G C  
 SEQ. ID. NO. 36 C A A G G C C A T G G C T G A G A T T C T C C G C  
 SEQ. ID. NO. 32 C C A G G C C A C T G C C A T G G C A G A C A T C

Figure 11c

SEQ. ID. NO. 40 T A C C A A G C C C A A G C C A T G G T G G A C A  
 SEQ. ID. NO. 46 T T C T T C A A C T G G A C C T A T G T G T C C A  
 SEQ. ID. NO. 36 T T C T T C A A C T G G A C C T A T G T G T C C A  
 SEQ. ID. NO. 32 A T C G A G T A T T T T C C G C T G G A A C T G G G

SEQ. ID. NO. 40 T C G T G A C A G C A C T G G G A T G G A A T T A  
 SEQ. ID. NO. 46 C T G T G G C G T C T G A G G G C G A C T A T G G  
 SEQ. ID. NO. 36 C T G T G G C G T C T G A G G G C G A C T A T G G  
 SEQ. ID. NO. 32 T G G G C A C A A T T G C A G C T G A T G A C G A

SEQ. ID. NO. 40 T G T T T C G A C A C T G G C T T C T G A G G G G  
 SEQ. ID. NO. 46 C G A G A C A G G C A T T G A G G C C T T T G A G  
 SEQ. ID. NO. 36 C G A G A C A G G C A T T G A G G C C T T T G A G  
 SEQ. ID. NO. 32 C T A T G G G C G G C C G G G G A T T G A G A A A

SEQ. ID. NO. 40 A A C T A T G G T G A G A G C G G T G T G G A G G  
 SEQ. ID. NO. 46 C T A G A G G C T C G T G C C C G C A A C A T C T  
 SEQ. ID. NO. 36 C T A G A G G C T C G T G C C C G C A A C A T C T  
 SEQ. ID. NO. 32 T T C C G A G A G G A A G C T G A G G A A A G G G

SEQ. ID. NO. 40 C C T T C A C C C A G A T C T C G A G G G A G A T  
 SEQ. ID. NO. 46 G T G T G G C C A C C T C G G A G A A A G T G G G  
 SEQ. ID. NO. 36 G T G T G G C C A C C T C G G A G A A A G T G G G  
 SEQ. ID. NO. 32 A T A T C T G C A T C G A C T T C A G T G A A C T

SEQ. ID. NO. 40 T G G T G G T G T T T G C A T T G C T C A G T C A  
 SEQ. ID. NO. 46 C C G T G C C A T G A G C C G C G C G G C C T T T  
 SEQ. ID. NO. 36 C C G T G C C A T G A G C C G C G C G G C C T T T  
 SEQ. ID. NO. 32 C A T C T C C C A G T A C T C T G A T G A G G A A

SEQ. ID. NO. 40 C A G A A A A T C C C A C G T G A A C C A A G A C  
 SEQ. ID. NO. 46 G A G G G T G T G G T G C G A G C C C T G C T G C  
 SEQ. ID. NO. 36 G A G G G T G T G G T G C G A G C C C T G C T G C  
 SEQ. ID. NO. 32 G A G A T C C A G C A T G T G G T A G A G G T G A

SEQ. ID. NO. 40 C T G G A G A A T T T G A A A A A A T T A T C A A  
 SEQ. ID. NO. 46 A G A A G C C C A G T G C C C G C G T G G C T G T  
 SEQ. ID. NO. 36 A G A A G C C C A G T G C C C G C G T G G C T G T  
 SEQ. ID. NO. 32 T T C A A A A T T C C A C G G C C A A A G T C A T

Figure 11d

SEQ. ID. NO. 40 A C G C C T G C T A G A A A C A C C T A A T G C T  
 SEQ. ID. NO. 46 C C T G T T C A C C C G T T C T G A G G A T G C C  
 SEQ. ID. NO. 36 C C T G T T C A C C C G T T C T G A G G A T G C C  
 SEQ. ID. NO. 32 C G T G G T T T T C T C A G T G G C C C A G A T

SEQ. ID. NO. 40 C G A G C A G T G A T T A T G T T T G C C A A T G  
 SEQ. ID. NO. 46 C G G G A G C T G C T T G C T G C C A G C C A G C  
 SEQ. ID. NO. 36 C G G G A G C T G C T T G C T G C C A G C C A G C  
 SEQ. ID. NO. 32 C T T G A G C C C C T C A T C A A G G A G A T T G

SEQ. ID. NO. 40 A G G A T G A C A T C A G G A G G A T A T T G G A  
 SEQ. ID. NO. 46 G C C T C A A T G C C A G C T T C A C C T G G G T  
 SEQ. ID. NO. 36 G C C T C A A T G C C A G C T T C A C C T G G G T  
 SEQ. ID. NO. 32 T C C G G C G C A A T A T C A C G G G C A A G A T

SEQ. ID. NO. 40 A G C A G C A A A A A A A C T A A A C C A A A G T  
 SEQ. ID. NO. 46 G G C C A G T G A T G G T T G G G G G G C C C T G  
 SEQ. ID. NO. 36 G G C C A G T G A T G G T T G G G G G G C C C T G  
 SEQ. ID. NO. 32 C T G G C T G G C C A G C G A G G C C T G G G C C

SEQ. ID. NO. 40 G G G C A T T T T C T C T G G A T T G G C T C A G  
 SEQ. ID. NO. 46 G A G A G T G T G G T G G C A G G C A G T G A G G  
 SEQ. ID. NO. 36 G A G A G T G T G G T G G C A G G C A G T G A G G  
 SEQ. ID. NO. 32 A G C T C C T C C C T G A T C G C C A T G C C T C

SEQ. ID. NO. 40 A T A G T T G G G G A T C C A A A A T A G C A C C  
 SEQ. ID. NO. 46 G G G C T G C T G A G G G T G C T A T C A C C A T  
 SEQ. ID. NO. 36 G G G C T G C T G A G G G T G C T A T C A C C A T  
 SEQ. ID. NO. 32 A G T A C T T C C A C G T G G T T G G C G G C A C

SEQ. ID. NO. 40 T G T C T A T C A G C A A G A G G A G A T T G C A  
 SEQ. ID. NO. 46 C G A G C T G G C C T C C T A C C C C A T C A G T  
 SEQ. ID. NO. 36 C G A G C T G G C C T C C T A C C C C A T C A G T  
 SEQ. ID. NO. 32 C A T T G G A T T C G C T C T G A A G G C T G G G

SEQ. ID. NO. 40 G A A G G G G C T G T G A C A A T T T T G C C C A  
 SEQ. ID. NO. 46 G A C T T T G C C T C C T A C T T C C A G A G C C  
 SEQ. ID. NO. 36 G A C T T T G C C T C C T A C T T C C A G A G C C  
 SEQ. ID. NO. 32 C A G A T C C C A G G C T T C C G G G A A T T C C

Figure 11e

SEQ. ID. NO. 40 A A C G A G C A T C A A T T G A T G G A T T T G A  
 SEQ. ID. NO. 46 T G G A C C C T T G G A A C A A C A G C C G G A A  
 SEQ. ID. NO. 36 T G G A C C C T T G G A A C A A C A G C C G G A A  
 SEQ. ID. NO. 32 T G A A G A A G G T C C A T C C C A G G A A G T C

SEQ. ID. NO. 40 T C G A T A C T T T A G A A G C C G A A C T C T T  
 SEQ. ID. NO. 46 C C C C T G G T T C C G T G A A T T C T G G G A G  
 SEQ. ID. NO. 36 C C C C T G G T T C C G T G A A T T C T G G G A G  
 SEQ. ID. NO. 32 T G T C C A C A A T G G T T T T G C C A A G G A G

SEQ. ID. NO. 40 G C C A A T A A T C G A A G A A A T G T G T G G T  
 SEQ. ID. NO. 46 C A G A G G T T C C G C T G C A G C T T C C G G C  
 SEQ. ID. NO. 36 C A G A G G T T C C G C T G C A G C T T C C G G C  
 SEQ. ID. NO. 32 T T T T G G G A A G A A A C A T T T A A C T G C C

SEQ. ID. NO. 40 T T G C A G A A T T C T G G G A G G A G A A T T T  
 SEQ. ID. NO. 46 A G C G A G A C T G C G C A G C C C A C T C T C T  
 SEQ. ID. NO. 36 A G C G A G A C T G C G C A G C C C A C T C T C T  
 SEQ. ID. NO. 32 A C C T C C A A G A A G G T G C A A A A G G A C C

SEQ. ID. NO. 40 T G G C T G C A A G T T A G G A T C A C A T G G G  
 SEQ. ID. NO. 46 C C G G G C T G T G C C C T T T G A G C A G G A G  
 SEQ. ID. NO. 36 C C G G G C T G T G C C C T T T G A G C A G G A G  
 SEQ. ID. NO. 32 T T T A C C T G T G G A C A C C T T T C T G A G A

SEQ. ID. NO. 40 A A A A G G A A C A G T C A T A T A A A G A A A T  
 SEQ. ID. NO. 46 T C C A A G A T C A T G T T T G T G G T C A A T G  
 SEQ. ID. NO. 36 T C C A A G A T C A T G T T T G T G G T C A A T G  
 SEQ. ID. NO. 32 G G T C A C G A A G A A A G T G G C G A C A G G T

SEQ. ID. NO. 40 G C A C A G G G C T G G A G C G A A T T G C T C G  
 SEQ. ID. NO. 46 C A G T G T A C G C C A T G G C C C A T G C G C T  
 SEQ. ID. NO. 36 C A G T G T A C G C C A T G G C C C A T G C G C T  
 SEQ. ID. NO. 32 T T A G C A A C A G C T C G A C A G C C T T C C G

SEQ. ID. NO. 40 G G A T T C A T C T T A T G A A C A G G A A G G A  
 SEQ. ID. NO. 46 C C A C A A C A T G C A C C G T G C C C T C T G C  
 SEQ. ID. NO. 36 C C A C A A C A T G C A C C G T G C C C T C T G C  
 SEQ. ID. NO. 32 A C C C C T C T G T A C A G G G G A T G A G A A C

Figure 11f

SEQ. ID. NO. 40 A A G G T C C A A T T T G T A A T T G A T G C T G  
 SEQ. ID. NO. 46 C C C A A C A C C A C C C G G C T C T G T G A C G  
 SEQ. ID. NO. 36 C C C A A C A C C A C C C G G C T C T G T G A C G  
 SEQ. ID. NO. 32 A T C A G C A G T G T C G A G A C C C C T T A C A

SEQ. ID. NO. 40 T A T A T T C C A T G G C T T A C G C C C T G C A  
 SEQ. ID. NO. 46 C G A T G C G G C C A G T T A A C G G G C G C C G  
 SEQ. ID. NO. 36 C G A T G C G G C C A G T T A A C G G G C G C C G  
 SEQ. ID. NO. 32 T A G A T T A C A C G C A T T T A C G G A T A T C

SEQ. ID. NO. 40 C A A T A T G C A C A A A G A T C T C T G C C C T  
 SEQ. ID. NO. 46 C C T C T A C A A G G A C T T T G T G C T C A A C  
 SEQ. ID. NO. 36 C C T C T A C A A G G A C T T T G T G C T C A A C  
 SEQ. ID. NO. 32 C T A C A A T G T G T A C T T A G C A G T C T A C

SEQ. ID. NO. 40 G G A T A C A T T G G C C T T T G T C C A C G A A  
 SEQ. ID. NO. 46 G T C A A G T T T G A T G C C C C C T T T C G C C  
 SEQ. ID. NO. 36 G T C A A G T T T G A T G C C C C C T T T C G C C  
 SEQ. ID. NO. 32 T C C A T T G C C C A C G C C T T G C A A G A T A

SEQ. ID. NO. 40 T G A G T A C C A T T G A T G G G A A A G A G C T  
 SEQ. ID. NO. 46 C A G C T G A C A C C C A C A A T G A G G T C C G  
 SEQ. ID. NO. 36 C A G C T G A C A C C C A C A A T G A G G T C C G  
 SEQ. ID. NO. 32 T A T A T A C C T G C T T A C C T G G G A G A G G

SEQ. ID. NO. 40 A C T T G G T T A T A T T C G G G C T G T A A A T  
 SEQ. ID. NO. 46 C T T T G A C C G C T T T G G T G A T G G T A T T  
 SEQ. ID. NO. 36 C T T T G A C C G C T T T G G T G A T G G T A T T  
 SEQ. ID. NO. 32 G C T C T T C A C C A A T G G C T C C T G T G C A

SEQ. ID. NO. 40 T T T A A T G G C A G T G C T G G C A C T C C T G  
 SEQ. ID. NO. 46 G G C C G C T A C A A C A T C T T C A C C T A T C  
 SEQ. ID. NO. 36 G G C C G C T A C A A C A T C T T C A C C T A T C  
 SEQ. ID. NO. 32 G A C A T C A A G A A A G T T G A G G C G T G G C

SEQ. ID. NO. 40 T C A C T T T T A A T G A A A A C G G A G A T G C  
 SEQ. ID. NO. 46 T G C G T G C A G G C A G T G G G C G C T A T C G  
 SEQ. ID. NO. 36 T G C G T G C A G G C A G T G G G C G C T A T C G  
 SEQ. ID. NO. 32 A G G T C C T G A A G C A C C T A C G G C A T C T

Figure 11g

SEQ. ID. NO. 40 TCCTGGACGTTATGATATCTTCCAG  
 SEQ. ID. NO. 46 CTACCAGAAAGGTGGGCTACTGGGCA  
 SEQ. ID. NO. 36 CTACCAGAAAGGTGGGCTACTGGGCA  
 SEQ. ID. NO. 32 AAAC TTTACAAACAATA TGGGGGAG

SEQ. ID. NO. 40 TATCAAATAACCAACAAAAGCACAG  
 SEQ. ID. NO. 46 GAAGGCTTTGACTCTGGACACCAGCC  
 SEQ. ID. NO. 36 GAAGGCTTTGACTCTGGACACCAGCC  
 SEQ. ID. NO. 32 CAGGTGACCTTTGATGAGTGTGGTG

SEQ. ID. NO. 40 AGTACAAAGTCAATCGGCCCACTGGAC  
 SEQ. ID. NO. 46 TCATCCCATGGGCCCTCACCCCTCAGC  
 SEQ. ID. NO. 36 TCATCCCATGGGCCCTCACCCCTCAGC  
 SEQ. ID. NO. 32 ACCGTGGTGGGGAAC TATTCCATCAT

SEQ. ID. NO. 40 CAATCAGCTTTCATCTAAAAGTGGAA  
 SEQ. ID. NO. 46 CGGCCCCCTTGCCCGCCTCTCGCTGC  
 SEQ. ID. NO. 36 CGGCCCCCTTGCCCGCCTCTCGCTGC  
 SEQ. ID. NO. 32 CAACTGGGCACTCTCCCCAGAGGAT

SEQ. ID. NO. 40 GACATGCAAGTGGGCTCATAGAGAAC  
 SEQ. ID. NO. 46 AGTGAGCCCTTGCCCTCCAGAAATGAGG  
 SEQ. ID. NO. 36 AGTGAGCCCTTGCCCTCCAGAAATGAGG  
 SEQ. ID. NO. 32 GGCTCCAATCGTGTTTTAAAGGAAGTCG

SEQ. ID. NO. 40 ATACTCACCCGGCGTCTGTCTGCAG  
 SEQ. ID. NO. 46 TGAAGAGGTGTGTCAGCCGGGCGAAGT  
 SEQ. ID. NO. 36 TGAAGAGGTGTGTCAGCCGGGCGAAGT  
 SEQ. ID. NO. 32 GGTATTACAAACGTCTATGCCAAGAA

SEQ. ID. NO. 40 CCTGCCGTTGTAAGCCAGGGGAGAGG  
 SEQ. ID. NO. 46 CTGCTGCTGGCTCTTGCA TTCCGTGC  
 SEQ. ID. NO. 36 CTGCTGCTGGCTCTTGCA TTCCGTGC  
 SEQ. ID. NO. 32 GGGAGAAAGACTCTTTCATCAACGAG

SEQ. ID. NO. 40 AAGAAAACGGTGAAAGGGGTCCCTT  
 SEQ. ID. NO. 46 CAGCCCTATGAGTACCGATTGGACG  
 SEQ. ID. NO. 36 CAGCCCTATGAGTACCGATTGGACG  
 SEQ. ID. NO. 32 GAGAAAATCCTGTGGAGTGGGTTCT

Figure 11h

SEQ. ID. NO. 40 G C T G C T G G C A C T G T G A A C G C T G T G A  
 SEQ. ID. NO. 46 A A T T C A C T T G C G C T G A T T G T G G C C T  
 SEQ. ID. NO. 36 A A T T C A C T T G C G C T G A T T G T G G C C T  
 SEQ. ID. NO. 32 C C A G G G A G G T G C C C T T C T C C A A C T G

SEQ. ID. NO. 40 A G G T T A C A A C T A C C A G G T G G A T G A G  
 SEQ. ID. NO. 46 G G G C T A C T G G C C C A A T G C C A G C C T G  
 SEQ. ID. NO. 36 G G G C T A C T G G C C C A A T G C C A G C C T G  
 SEQ. ID. NO. 32 C A G C C G A G A C T G C C T G G C A G G G A C C

SEQ. ID. NO. 40 C T G T C C T G T G A A C T T T G C C C T C T G G  
 SEQ. ID. NO. 46 A C T G G C T G C T T C G A A C T G C C C C A G G  
 SEQ. ID. NO. 36 A C T G G C T G C T T C G A A C T G C C C C A G G  
 SEQ. ID. NO. 32 A G G A A A G G G A T C A T T G A G G G G G A G C

SEQ. ID. NO. 40 A T C A G A G A C C C A A C A T G A A C C G C A C  
 SEQ. ID. NO. 46 A G T A C A T C C G C T G G G G C G A T G C C T G  
 SEQ. ID. NO. 36 A G T A C A T C C G C T G G G G C G A T G C C T G  
 SEQ. ID. NO. 32 C C A C C T G C T G C T T T G A G T G T G T G G A

SEQ. ID. NO. 40 A G G C T G C C A G C T T A T C C C C A T C A T C  
 SEQ. ID. NO. 46 G G C T G T G G G A C C T G T C A C C A T C G C C  
 SEQ. ID. NO. 36 G G C T G T G G G A C C T G T C A C C A T C G C C  
 SEQ. ID. NO. 32 G T G T C C T G A T G G G G A G T A T A G T G A T

SEQ. ID. NO. 40 A A A T T G G A G T G G C A T T C T C C C T G G G  
 SEQ. ID. NO. 46 T G C C T C G G T G C C C T G G C C A C C C T C T  
 SEQ. ID. NO. 36 T G C C T C G G T G C C C T G G C C A C C C T C T  
 SEQ. ID. NO. 32 G A G A C A G A T G C C A G T G C C T G T A A C A

SEQ. ID. NO. 40 C T G T G G T G C C T G T G T T T G T T G C A A T  
 SEQ. ID. NO. 46 T T G T G C T G G G T G T C T T T G T G C G G C A  
 SEQ. ID. NO. 36 T T G T G C T G G G T G T C T T T G T G C G G C A  
 SEQ. ID. NO. 32 A G T G C C C A G A T G A C T T C T G G T C C A A

SEQ. ID. NO. 40 A T T G G G A A T C A T C G C C A C C A C C T T T  
 SEQ. ID. NO. 46 C A A T G C C A C A C C A G T G G T C A A G G C C  
 SEQ. ID. NO. 36 C A A T G C C A C A C C A G T G G T C A A G G C C  
 SEQ. ID. NO. 32 T G A G A A C C A C A C C T C C T G C T T C G A A

Figure 11i



SEQ. ID. NO. 40 GTGATCGTGACCTTTGTCCGCTATA  
 SEQ. ID. NO. 46 TCAGGTCGGGAGCTCTGCTACATCC  
 SEQ. ID. NO. 36 TCAGGTCGGGAGCTCTGCTACATCC  
 SEQ. ID. NO. 32 CTGCCCCAGGAGTACATCCGCTGGG

SEQ. ID. NO. 40 ATGACACACCTATCGTGAGGGCTTC  
 SEQ. ID. NO. 46 TGCTGGGTGGTGTCTTCCTCTGCTA  
 SEQ. ID. NO. 36 TGCTGGGTGGTGTCTTCCTCTGCTA  
 SEQ. ID. NO. 32 GCGATGCCCTGGGCTGTGGGACCTGT

SEQ. ID. NO. 40 AGGACGCGAACTTAGTTACGTGCTC  
 SEQ. ID. NO. 46 CTGCATGACCTTTCATCTTCATTGCC  
 SEQ. ID. NO. 36 CTGCATGACCTTTCATCTTCATTGCC  
 SEQ. ID. NO. 32 CACCATCGCCCTGCCCTCGGTGCCCTG

SEQ. ID. NO. 40 CTAACGGGGGATTTTCTCTGTTATT  
 SEQ. ID. NO. 46 AAGCCATCCACGGCAGTGTTACCT  
 SEQ. ID. NO. 36 AAGCCATCCACGGCAGTGTTACCT  
 SEQ. ID. NO. 32 GCCACCCTGTTTGTGCTGGGTGTCT

SEQ. ID. NO. 40 CAATCACGTTTTTTAATGATTGCAAGC  
 SEQ. ID. NO. 46 TACGGCGTCTTTGGTTTGGGGCACTGC  
 SEQ. ID. NO. 36 TACGGCGTCTTTGGTTTGGGGCACTGC  
 SEQ. ID. NO. 32 TTGTGCGGCAACAATGCCACACCAGT

SEQ. ID. NO. 40 ACCAGATACAATCATATGCTCCTTTC  
 SEQ. ID. NO. 46 CTTCTCTGTCTGTCTACTCAGCCCTG  
 SEQ. ID. NO. 36 CTTCTCTGTCTGTCTACTCAGCCCTG  
 SEQ. ID. NO. 32 GGTC AAGGCCCTCAGGTCGGGAGCTC

SEQ. ID. NO. 40 CGACGGGGTCTTTCCTAGGACTTGGCA  
 SEQ. ID. NO. 46 CTCACCAAGACCAACC GCATTGCAC  
 SEQ. ID. NO. 36 CTCACCAAGACCAACC GCATTGCAC  
 SEQ. ID. NO. 32 TGCTACATCCTGCTGGGTGGTGTCT

SEQ. ID. NO. 40 TGTGTTTTCAGCTATGCAAGCCCTTCT  
 SEQ. ID. NO. 46 GCATCTTTCGGTGGGGGCCCGGGAGGG  
 SEQ. ID. NO. 36 GCATCTTTCGGTGGGGGCCCGGGAGGG  
 SEQ. ID. NO. 32 TCTCTGTGCTACTGCATGACCTTCAAT

Figure 11j

SEQ. ID. NO. 40 G A C C A A A A C A A A C C G T A T C C A C C G A  
 SEQ. ID. NO. 46 T G C C C A G C G G C C A C G C T T C A T C A G T  
 SEQ. ID. NO. 36 T G C C C A G C G G C C A C G C T T C A T C A G T  
 SEQ. ID. NO. 32 C T T C A T T G C C A A G C C A T C C A C G G C A

SEQ. ID. NO. 40 A T A T T T G A G C A G G G G A A G A A A T C T G  
 SEQ. ID. NO. 46 C C T G C C T C A C A G G T G G C C A T C T G C C  
 SEQ. ID. NO. 36 C C T G C C T C A C A G G T G G C C A T C T G C C  
 SEQ. ID. NO. 32 G T G T G T A C C T T A C G G C G T C T T G G T T

SEQ. ID. NO. 40 T C A C A G C G C C C A A G T T C A T T A G T C C  
 SEQ. ID. NO. 46 T G G C A C T T A T C T C G G G C C A G C T G C T  
 SEQ. ID. NO. 36 T G G C A C T T A T C T C G G G C C A G C T G C T  
 SEQ. ID. NO. 32 T G G G C A C T G C C T T C T C T G T C T G C T A

SEQ. ID. NO. 40 A G C A T C T C A G C T G G T G A T C A C C T T C  
 SEQ. ID. NO. 46 C A T C G T G G T C G C C T G G C T G G T G G T G  
 SEQ. ID. NO. 36 C A T C G T G G T C G C C T G G C T G G T G G T G  
 SEQ. ID. NO. 32 C T C A G C C C T G C T C A C C A A G A C C A A C

SEQ. ID. NO. 40 A G C C T C A T C T C C G T C C A G C T C C T T G  
 SEQ. ID. NO. 46 G A G G C A C C G G G C A C A G G C A A G G A G A  
 SEQ. ID. NO. 36 G A G G C A C C G G G C A C A G G C A A G G A G A  
 SEQ. ID. NO. 32 C G C A T T G C A C G C A T C T T C G G T G G G G

SEQ. ID. NO. 40 G A G T G T T T G T C T G G T T T G T T G T G G A  
 SEQ. ID. NO. 46 C A G C C C C C G A A C G G C G G G A G G T G G T  
 SEQ. ID. NO. 36 C A G C C C C C G A A C G G C G G G A G G T G G T  
 SEQ. ID. NO. 32 C C C G G G A G G G T G C C C A G C G G C C A C G

SEQ. ID. NO. 40 T C C C C C C C A C A T C A T C A T T G A C T A T  
 SEQ. ID. NO. 46 G A C A C T G C G C T G C A A C C A C C G C G A T  
 SEQ. ID. NO. 36 G A C A C T G C G C T G C A A C C A C C G C G A T  
 SEQ. ID. NO. 32 C T T C A T C A G T C C T G C C T C A C A G G T G

SEQ. ID. NO. 40 G G A G A G C A G C G G A C A C T A G A T C C A G  
 SEQ. ID. NO. 46 G C A A G T A T G T T G G G C T C G C T G G C C T  
 SEQ. ID. NO. 36 G C A A G T A T G T T G G G C T C G C T G G C C T  
 SEQ. ID. NO. 32 G C C A T C T G C C T G G C A C T T A T C T C G G

Figure 11k

SEQ. ID. NO. 40 A G A A G G C C A G G G G A G T G C T C A A G T G  
 SEQ. ID. NO. 46 A C A A T G T G C T C C T C A T C G C G C T C T G  
 SEQ. ID. NO. 36 A C A A T G T G C T C C T C A T C G C G C T C T G  
 SEQ. ID. NO. 32 G C C A G C T G C T C A T C G T G G T C G C C T G

SEQ. ID. NO. 40 T G A C A T T T C T G A T C T C T C A C T C A T T  
 SEQ. ID. NO. 46 C A C G C T T T A T G C C T T C A A G A C T C G C  
 SEQ. ID. NO. 36 C A C G C T T T A T G C C T T C A A G A C T C G C  
 SEQ. ID. NO. 32 G C T G G T G G T G G A G G C A C C G G G C A C A

SEQ. ID. NO. 40 T G T T C A C T T G G A T A C A G T A T C C T C T  
 SEQ. ID. NO. 46 A A G T G C C C C G A A A A C T T C A A C G A G G  
 SEQ. ID. NO. 36 A A G T G C C C C G A A A A C T T C A A C G A G G  
 SEQ. ID. NO. 32 G G C A A G G A G A C A G C C C C C G A A C G G C

SEQ. ID. NO. 40 T G A T G G T C A C T T G T A C T G T T T A T G C  
 SEQ. ID. NO. 46 C C A A G T T C A T T G G C T T C A C C A T G T A  
 SEQ. ID. NO. 36 C C A A G T T C A T T G G C T T C A C C A T G T A  
 SEQ. ID. NO. 32 G G G A G G T G G T G A C A C T G C G C T G C A A

SEQ. ID. NO. 40 C A T T A A A A C G A G A G G T G T C C C A G A G  
 SEQ. ID. NO. 46 C A C C A C C T G C A T C A T C T G G C T G G C A  
 SEQ. ID. NO. 36 C A C C A C C T G C A T C A T C T G G C T G G C A  
 SEQ. ID. NO. 32 C C A C C G C G A T G C A A G T A T G T T G G G C

SEQ. ID. NO. 40 A C T T T C A A T G A A G C C A A A C C T A T T G  
 SEQ. ID. NO. 46 T T C C T G C C C A T C T T C T A T G T C A C C T  
 SEQ. ID. NO. 36 T T C C T G C C C A T C T T C T A T G T C A C C T  
 SEQ. ID. NO. 32 T C G C T G G C C T A C A A T G T G C T C C T C A

SEQ. ID. NO. 40 G A T T T A C C A T G T A T A C C A C C T G C A T  
 SEQ. ID. NO. 46 C C A G T G A C T A C C G G G T A C A G A C C A C  
 SEQ. ID. NO. 36 C C A G T G A C T A C C G G G T A C A G A C C A C  
 SEQ. ID. NO. 32 T C G C G C T C T G C A C G C T T T A T G C C T T

SEQ. ID. NO. 40 C A T T T G G T T A G C T T T C A T C C C C A T C  
 SEQ. ID. NO. 46 C A C C A T G T G C G T G T C A G T C A G C C T C  
 SEQ. ID. NO. 36 C A C C A T G T G C G T G T C A G T C A G C C T C  
 SEQ. ID. NO. 32 C A A T A C T C G C A A G T G C C C C G A A A A C

Figure 11L

SEQ. ID. NO. 40 T T T T T T G G T A C A G C C C A G T C A G C A G  
 SEQ. ID. NO. 46 A G C G G C T C C G T G G T G C T T G G C T G C C  
 SEQ. ID. NO. 36 A G C G G C T C C G T G G T G C T T G G C T G C C  
 SEQ. ID. NO. 32 T T C A A C G A G G C C A A G T T C A T T G G C T

SEQ. ID. NO. 40 A A A A G A T G T A C A T C C A G A C A A C A A C  
 SEQ. ID. NO. 46 T C T T T G C G C C C A A G C T G C A C A T C A T  
 SEQ. ID. NO. 36 T C T T T G C G C C C A A G C T G C A C A T C A T  
 SEQ. ID. NO. 32 T C A C C A T G T A C A C C A C C T G C A T C A T

SEQ. ID. NO. 40 A C T T A C T G T C T C C A T G A G T T T A A G T  
 SEQ. ID. NO. 46 C C T C T T C C A G C C G C A G A A G A A C A C C  
 SEQ. ID. NO. 36 C C T C T T C C A G C C G C A G A A G A A C A C C  
 SEQ. ID. NO. 32 C T G G C T G G C A T T G T T G C C C A T C T T C

SEQ. ID. NO. 40 G C T T C A G T A T C T C T G G G C A T G C T C T  
 SEQ. ID. NO. 46 A T C G A G G A G G T G C G T T G C A G C A C C G  
 SEQ. ID. NO. 36 A T C G A G G A G G T G C G T T G C A G C A C C G  
 SEQ. ID. NO. 32 T A T G T C A C C T C C A G T G A C T A C C G G G

SEQ. ID. NO. 40 A T A T G C C C A A G G T T T A T A T T A T A A T  
 SEQ. ID. NO. 46 C A G C T C A C G C T T T C A A G G T G G C T G C  
 SEQ. ID. NO. 36 C A G C T C A C G C T T T C A A G G T G G C T G C  
 SEQ. ID. NO. 32 T A C A G A C C A C C A C C A T G T G C G T G T C

SEQ. ID. NO. 40 T T T T C A T C C A G A A C A G A A T A C C A T C  
 SEQ. ID. NO. 46 C C G G G C C A C G C T G C G C C G C A G C A A C  
 SEQ. ID. NO. 36 C C G G G C C A C G C T G C G C C G C A G C A A C  
 SEQ. ID. NO. 32 A G T C A G C C T C A G C G G C T C C G T G G T G

SEQ. ID. NO. 40 G A G G A G G T G C G T T G C A G C A C C G C A G  
 SEQ. ID. NO. 46 G T C T C C C G C A A G C G G T C C A G C A G C C  
 SEQ. ID. NO. 36 G T C T C C C G C A A G C G G T C C A G C A G C C  
 SEQ. ID. NO. 32 C T T G G C T G C C T C T T T G C G C C C A A G C

SEQ. ID. NO. 40 C T C A C G C T T T C A A G G T G G C T G C C C G  
 SEQ. ID. NO. 46 T T G G A G G C T C C A C G G G A T C C A C C C C  
 SEQ. ID. NO. 36 T T G G A G G C T C C A C G G G A T C C A C C C C  
 SEQ. ID. NO. 32 T G C A C A T C A T C C T C T T C C A G C C G C A

Figure 11m

SEQ. ID. NO. 40 G G C C A C G C T G C G C C G C A G C A A C G T C  
 SEQ. ID. NO. 46 C T C C T C C T C C A T C A G C A G C A A G A G C  
 SEQ. ID. NO. 36 C T C C T C C T C C A T C A G C A G C A A G A G C  
 SEQ. ID. NO. 32 G A A G A A C G T G G T T A G C C A C C G G G C A

SEQ. ID. NO. 40 T C C C G C A A G C G G T C C A G C A G C C T T G  
 SEQ. ID. NO. 46 A A C A G C G A A G A C C C A T T C C C A C A G C  
 SEQ. ID. NO. 36 A A C A G C G A A G A C C C A T T C C C A C A G C  
 SEQ. ID. NO. 32 C C C A C C A G C C G C T T T G G C A G T G C T G

SEQ. ID. NO. 40 G A G G C T C C A C G G G A T C C A C C C C C T C  
 SEQ. ID. NO. 46 C C G A G A G G C A G A A A G C A G C A G C A G C C  
 SEQ. ID. NO. 36 C C G A G A G G C A G A A A G C A G C A G C A G C C  
 SEQ. ID. NO. 32 C T G C C A G G G C C A G C T C C A G C C T T G G

SEQ. ID. NO. 40 C T C C T C C A T C A G C A G C A A G A G C A A C  
 SEQ. ID. NO. 46 G C T G G C C C T A A C C C A G C A A G A G C A G  
 SEQ. ID. NO. 36 G C T G G C C C T A A C C C A G C A A G A G C A G  
 SEQ. ID. NO. 32 C C A A G G G T C T G G C T C C C A G T T T G T C

SEQ. ID. NO. 40 A G C G A A G A C C C A T T C C C A C A G C C C G  
 SEQ. ID. NO. 46 C A G C A G C A G C C C C T G A C C C T C C C A C  
 SEQ. ID. NO. 36 C A G C A G C A G C C C C T G A C C C T C C C A C  
 SEQ. ID. NO. 32 C C C A C T G T T T G C A A T G G C C G T G A G G

SEQ. ID. NO. 40 A G A G G C A G A A A G C A G C A G C A G C C G C T  
 SEQ. ID. NO. 46 A G C A G C A A C G A T C T C A G C A G C A G C C  
 SEQ. ID. NO. 36 A G C A G C A A C G A T C T C A G C A G C A G C C  
 SEQ. ID. NO. 32 T G G T G G A C T C G A C A A A C G T C A T C G C T

SEQ. ID. NO. 40 G G C C C T A A C C C A G C A A A G A G C A G C A G  
 SEQ. ID. NO. 46 C A G A T G C A A A G C A G A A A G G T C A T C T T T  
 SEQ. ID. NO. 36 C A G A T G C A A A G C A G A A A G G T C A T C T T T  
 SEQ. ID. NO. 32 T A T G A C T C T G G A G T C C A T C A T G G C G

SEQ. ID. NO. 40 C A G C A G C C C C T G A C C C T C C C A C A G C  
 SEQ. ID. NO. 46 G G C A G C G G C A C G G T C A C C T T C T C A C  
 SEQ. ID. NO. 36 G G C A G C G G C A C G G T C A C C T T C T C A C  
 SEQ. ID. NO. 32 T G C T G C C T G A G C G A G G A G G C C A A G G

Figure 11n

SEQ. ID. NO. 40 A G C A A C G A T C T C A G C A G C A G C C C A G  
 SEQ. ID. NO. 46 T G A G C T T T G A T G A G C C T C A G A A G A A  
 SEQ. ID. NO. 36 T G A G C T T T G A T G A G C C T C A G A A G A A  
 SEQ. ID. NO. 32 A A G C C C G G C G G A T C A A C G A C G A G A T

SEQ. ID. NO. 40 A T G C A A G C A G A A G G T C A T C T T T G G C  
 SEQ. ID. NO. 46 C G C C A T G G C C C A C G G G A A T T C T A C G  
 SEQ. ID. NO. 36 C G C C A T G G C C C A C G G G A A T T C T A C G  
 SEQ. ID. NO. 32 C G A G C G G C A G C T C C G C A G G G A C A A G

SEQ. ID. NO. 40 A G C G G C A C G G T C A C C T T C T C A C T G A  
 SEQ. ID. NO. 46 C A C C A G A A C T C C C T G G A G G C C C A G A  
 SEQ. ID. NO. 36 C A C C A G A A C T C C C T G G A G G C C C A G A  
 SEQ. ID. NO. 32 C G G G A C G C C C G C C G G G A G C T C A A G C

SEQ. ID. NO. 40 G C T T T G A T G A G C C T C A G A A G A A C G C  
 SEQ. ID. NO. 46 A A A G C A G C G A T A C G C T G A C C C G A C A  
 SEQ. ID. NO. 36 A A A G C A G C G A T A C G C T G A C C C G A C A  
 SEQ. ID. NO. 32 T G C T G C T G C T C G G G A C A G G A G A G A G

SEQ. ID. NO. 40 C A T G G C C C A C G G G A A T T C T A C G C A C  
 SEQ. ID. NO. 46 C C A G C C A T T A C T C C C G C T G C A G T G C  
 SEQ. ID. NO. 36 C C A G C C A T T A C T C C C G C T G C A G T G C  
 SEQ. ID. NO. 32 T G G C A A G A G T A C G T T T A T C A A G C A G

SEQ. ID. NO. 40 C A G A A C T C C C T G G A G G C C C A G A A A A  
 SEQ. ID. NO. 46 G G G G A A A C G G A C T T A G A T C T G A C C G  
 SEQ. ID. NO. 36 G G G G A A A C G G A C T T A G A T C T G A C C G  
 SEQ. ID. NO. 32 A T G A G A A T C A T C C A T G G G T C A G G A T

SEQ. ID. NO. 40 G C A G C G A T A C G C T G A C C C G A C A C C A  
 SEQ. ID. NO. 46 T C C A G G A A A C A G G T C T G C A A G G A C C  
 SEQ. ID. NO. 36 T C C A G G A A A C A G G T C T G C A A G G A C C  
 SEQ. ID. NO. 32 A C T C T G A T G A A G A T A A A A G G G G C T T

SEQ. ID. NO. 40 G C C A T T A C T C C C G C T G C A G T G C G G G  
 SEQ. ID. NO. 46 T G T G G G T G G A G A C C A G C G G C C A G A G  
 SEQ. ID. NO. 36 T G T G G G T G G A G A C C A G C G G C C A G A G  
 SEQ. ID. NO. 32 C A C C A A G C T G G T G T A T C A G A A C A T C

Figure 110

SEQ. ID. NO. 40 G A A A C G G A C T T A G A T C T G A C C G T C C  
 SEQ. ID. NO. 46 G T G G A G G A C C C T G A A G A G T T G T C C C  
 SEQ. ID. NO. 36 G T G G A G G A C C C T G A A G A G T T G T C C C  
 SEQ. ID. NO. 32 T T C A C G G C C A T G C A G G C C A T G A T C A

SEQ. ID. NO. 40 A G G A A A C A G G T C T G C A A G G A C C T G T  
 SEQ. ID. NO. 46 C A G C A C T T G T A G T G T C C A G T T C A C A  
 SEQ. ID. NO. 36 C A G C A C T T G T A G T G T C C A G T T C A C A  
 SEQ. ID. NO. 32 G A G C C A T G G A C A C A C T C A A G A T C C C

SEQ. ID. NO. 40 G G G T G G A G A C C A G C G G C C A G A G G T G  
 SEQ. ID. NO. 46 G A G C T T T G T C A T C A G T G G T G G A G G C  
 SEQ. ID. NO. 36 G A G C T T T G T C A T C A G T G G T G G A G G C  
 SEQ. ID. NO. 32 A T A C A A G T A T G A G C A C A A T A A G G C T

SEQ. ID. NO. 40 G A G G A C C C T G A A G A G T T G T C C C C A G  
 SEQ. ID. NO. 46 A G C A C T G T T A C A G A A A A C G T A G T G A  
 SEQ. ID. NO. 36 A G C A C T G T T A C A G A A A A C G T A G T G A  
 SEQ. ID. NO. 32 C A T G C A C A A T T A G T T C G A G A A G T T G

SEQ. ID. NO. 40 C A C T T G T A G T G T C C A G T T C A C A G A G  
 SEQ. ID. NO. 46 A T T C A G C G G C C G C C A T G A C T C T G G A  
 SEQ. ID. NO. 36 A T T C A A T G A C T C T G G A G T C C A T C A T  
 SEQ. ID. NO. 32 A T G T G G A G A A G G T G T C T G C T T T T G A

SEQ. ID. NO. 40 C T T T G T C A T C A G T G G T G G A G G C A G C  
 SEQ. ID. NO. 46 G T C C A T C A T G G C G T G C T G C C T G A G C  
 SEQ. ID. NO. 36 G G C G T G C T G C C T G A G C G A G G A G G C C  
 SEQ. ID. NO. 32 G A A T C C A T A T G T A G A T G C A A T A A A G

SEQ. ID. NO. 40 A C T G T T A C A G A A A A C G T A G T G A A T T  
 SEQ. ID. NO. 46 G A G G A G G C C A A G G A A G C C C G G C G G A  
 SEQ. ID. NO. 36 A A G G A A G C C C G G C G G A T C A A C G A C G  
 SEQ. ID. NO. 32 A G T T T A T G G A A T G A T C C T G G A A T C C

SEQ. ID. NO. 40 C A - - - - -  
 SEQ. ID. NO. 46 T C A A C G A C G A G A T C G A G C G G C A G C T  
 SEQ. ID. NO. 36 A G A T C G A G C G G C A G C T C C G C A G G G A  
 SEQ. ID. NO. 32 A G G A A T G C T A T G A T A G A C G A C G A G A

Figure 11p

SEQ. ID. NO. 40 - - - - -  
 SEQ. ID. NO. 46 C C G C A G G G A C A A G C G G G A C G C C C G C  
 SEQ. ID. NO. 36 C A A G C G G G A C G C C C G C C G G G A G C T C  
 SEQ. ID. NO. 32 A T A T C A A T T A T C T G A C T C T A C C A A A

SEQ. ID. NO. 40 - - - - - A T G A C T C T G G  
 SEQ. ID. NO. 46 C G G G A G C T C A A G C T G C T G C T G C T C G  
 SEQ. ID. NO. 36 A A G C T G C T G C T G C T C G G G A C A G G A G  
 SEQ. ID. NO. 32 T A C T A T C T T A A T G A C T T G G A C C G C G

SEQ. ID. NO. 40 A G T C C A T C A T G G C G T G C T G C C T G A G  
 SEQ. ID. NO. 46 G G A C A G G A G A G A G T G G C A A G A G T A C  
 SEQ. ID. NO. 36 A G A G T G G C A A G A G T A C G T T T A T C A A  
 SEQ. ID. NO. 32 T A G C T G A C C C T G C C T A C C T G C C T A C

SEQ. ID. NO. 40 C G A G G A G G C C A A G G A A G C C C G G C G G  
 SEQ. ID. NO. 46 G T T T A T C A A G C A G A T G A G A A T C A T C  
 SEQ. ID. NO. 36 G C A G A T G A G A A T C A T C C A T G G G T C A  
 SEQ. ID. NO. 32 G C A A C A A G A T G T G C T T A G A G T T C G A

SEQ. ID. NO. 40 A T C A A C G A C G A G A T C G A G C G G C A G C  
 SEQ. ID. NO. 46 C A T G G G T C A G G A T A C T C T G A T G A A G  
 SEQ. ID. NO. 36 G G A T A C T C T G A T G A A G A T A A A A G G G  
 SEQ. ID. NO. 32 G T C C C C A C C A C A G G G A T C A T C G A A T

SEQ. ID. NO. 40 T C C G C A G G G A C A A G C G G G A C G C C C G  
 SEQ. ID. NO. 46 A T A A A A G G G G C T T C A C C A A G C T G G T  
 SEQ. ID. NO. 36 G C T T C A C C A A G C T G G T G T A T C A G A A  
 SEQ. ID. NO. 32 A C C C C T T T G A C T T A C A A A G T G T C A T

SEQ. ID. NO. 40 C C G G G A G C T C A A G C T G C T G C T G C T C  
 SEQ. ID. NO. 46 G T A T C A G A A C A T C T T C A C G G C C A T G  
 SEQ. ID. NO. 36 C A T C T T C A C G G C C A T G C A G G C C A T G  
 SEQ. ID. NO. 32 T T T C A G A A T G G T C G A T G T A G G G G G C

SEQ. ID. NO. 40 G G G A C A G G A G A G A G T G G C A A G A G T A  
 SEQ. ID. NO. 46 C A G G C C A T G A T C A G A G C C A T G G A C A  
 SEQ. ID. NO. 36 A T C A G A G C C A T G G A C A C A C T C A A G A  
 SEQ. ID. NO. 32 C A A A G G T C A G A G A G A A G A A A A T G G A

Figure 11q



SEQ. ID. NO. 40 C G T T T A T C A A G C A G A T G A G A A T C A T  
 SEQ. ID. NO. 46 C A C T C A A G A T C C C A T A C A A G T A T G A  
 SEQ. ID. NO. 36 T C C C A T A C A A G T A T G A G C A C A A T A A  
 SEQ. ID. NO. 32 T A C A C T G C T T T G A A A A T G T C A C C T C

SEQ. ID. NO. 40 C C A T G G G T C A G G A T A C T C T G A T G A A  
 SEQ. ID. NO. 46 G C A C A A T A A G G C T C A T G C A C A A T T A  
 SEQ. ID. NO. 36 G G C T C A T G C A C A A T T A G T T C G A G A A  
 SEQ. ID. NO. 32 T A T C A T G T T T C T A G T A G C G C T T A G T

SEQ. ID. NO. 40 G A T A A A A G G G G C T T C A C C A A G C T G G  
 SEQ. ID. NO. 46 G T T C G A G A A G T T G A T G T G G A G A A G G  
 SEQ. ID. NO. 36 G T T G A T G T G G A G A A G G T G T C T G C T T  
 SEQ. ID. NO. 32 G A A T A T G A T C A A G T T C T C G T G G A G T

SEQ. ID. NO. 40 T G T A T C A G A A C A T C T T C A C G G C C A T  
 SEQ. ID. NO. 46 T G T C T G C T T T T G A G A A T C C A T A T G T  
 SEQ. ID. NO. 36 T T G A G A A T C C A T A T G T A G A T G C A A T  
 SEQ. ID. NO. 32 C A G A C A A T G A G A A C C G A A T G G A G G A

SEQ. ID. NO. 40 G C A G G C C A T G A T C A G A G C C A T G G A C  
 SEQ. ID. NO. 46 A G A T G C A A T A A A G A G T T T A T G G A A T  
 SEQ. ID. NO. 36 A A A G A G T T T A T G G A A T G A T C C T G G A  
 SEQ. ID. NO. 32 A A G C A A G G C T C T C T T T A G A A C A A T T

SEQ. ID. NO. 40 A C A C T C A A G A T C C C A T A C A A G T A T G  
 SEQ. ID. NO. 46 G A T C C T G G A A T C C A G G A A T G C T A T G  
 SEQ. ID. NO. 36 A T C C A G G A A T G C T A T G A T A G A C G A C  
 SEQ. ID. NO. 32 A T C A C A T A C C C C T G G T T C C A G A A C T

SEQ. ID. NO. 40 A G C A C A A T A A G G C T C A T G C A C A A T T  
 SEQ. ID. NO. 46 A T A G A C G A C G A G A A T A T C A A T T A T C  
 SEQ. ID. NO. 36 G A G A A T A T C A A T T A T C T G A C T C T A C  
 SEQ. ID. NO. 32 C C T C G G T T A T T C T G T T C T T A A A C A A

SEQ. ID. NO. 40 A G T T C G A G A A G T T G A T G T G G A G A A G  
 SEQ. ID. NO. 46 T G A C T C T A C C A A A T A C T A T C T T A A T  
 SEQ. ID. NO. 36 C A A A T A C T A T C T T A A T G A C T T G G A C  
 SEQ. ID. NO. 32 G A A A G A T C T T C T A G A G G A G A A A A T C

Figure 11r

SEQ. ID. NO. 40 GTGTCCTGCTTTTGTAGGAATCCATATG  
 SEQ. ID. NO. 46 GACTTTGGACCGCGTAGCTGACCCCTG  
 SEQ. ID. NO. 36 CGCGTAGCTGACCCCTGCCTACCTGC  
 SEQ. ID. NO. 32 ATGTATTCCCATCTAGTCGACTACT

SEQ. ID. NO. 40 TAGATGCAATAAAGAGTTTATGGAA  
 SEQ. ID. NO. 46 CCTACCTGCCTACGCAACAAGATGT  
 SEQ. ID. NO. 36 CTACGCAACAAGATGTGCTTAGAGT  
 SEQ. ID. NO. 32 TCCCAGAAATATGATGGACCCCAAGAG

SEQ. ID. NO. 40 TGATCCTGGAATCCAGGAATGCTAT  
 SEQ. ID. NO. 46 GCTTAGAGTTTCGAGTCCCCACCCACA  
 SEQ. ID. NO. 36 TCGAGTCCCCACACAGGGATCATC  
 SEQ. ID. NO. 32 AGATGCCCAAGGCAGCCCCGAGAAATTC

SEQ. ID. NO. 40 GATAGACGACGAGAAATATCAATTAT  
 SEQ. ID. NO. 46 GGGATCATCGAATAACCCCTTTGACT  
 SEQ. ID. NO. 36 GAATAACCCCTTTGACTTACAAAGTG  
 SEQ. ID. NO. 32 ATTCTGAAGATGTTTCGTGGACCTGA

SEQ. ID. NO. 40 CTGACTCTACCAAAATACTATCTTAA  
 SEQ. ID. NO. 46 TACAAAGTGTCATTTTTCAGAAATGGT  
 SEQ. ID. NO. 36 TCATTTTTCAGAAATGGTCGATGTAGG  
 SEQ. ID. NO. 32 ACCCAGACAGTGACAAAAATTATCTA

SEQ. ID. NO. 40 TGACTTTGGACCGCGTAGCTGACCCCT  
 SEQ. ID. NO. 46 CGATGTAGGGGGGCCAAAGGTCAGAG  
 SEQ. ID. NO. 36 GGGCCAAAGGTCAGAGAGAGAGAAAAA  
 SEQ. ID. NO. 32 CTC CCACTTCAACGTGCGCCACAGAC

SEQ. ID. NO. 40 GCCTACCTGCCTACGCAACAAGATG  
 SEQ. ID. NO. 46 AGAAGAAAAATGGATACACTGCTTTTG  
 SEQ. ID. NO. 36 TGGATACACTGCTTTTGAAAAATGTCA  
 SEQ. ID. NO. 32 ACCGAGAAATATCCGCTTTTGTC TTG

SEQ. ID. NO. 40 TGCTTAGAGTTTCGAGTCCCCACCCAC  
 SEQ. ID. NO. 46 AAAATGTCACTCTATCATGTTTCT  
 SEQ. ID. NO. 36 CCTCTATCATGTTTCTAGTAGCGCT  
 SEQ. ID. NO. 32 CTGCCGTCAAGGACACCATCCTCCA

Figure 11s

SEQ. ID. NO. 40 A G G G A T C A T C G A A T A C C C C T T T G A C  
 SEQ. ID. NO. 46 A G T A G C G C T T A G T G A A T A T G A T C A A  
 SEQ. ID. NO. 36 T A G T G A A T A T G A T C A A G T T C T C G T G  
 SEQ. ID. NO. 32 G T T G A A C C T G A A G G A C T G C G G T C T G

SEQ. ID. NO. 40 T T A C A A A G T G T C A T T T T C A G A A T G G  
 SEQ. ID. NO. 46 G T T C T C G T G G A G T C A G A C A A T G A G A  
 SEQ. ID. NO. 36 G A G T C A G A C A A T G A G A A C C G A A T G G  
 SEQ. ID. NO. 32 T T C T A A

SEQ. ID. NO. 40 T C G A T G T A G G G G G C C A A A G G T C A G A  
 SEQ. ID. NO. 46 A C C G A A T G G A G G A A A G C A A G G C T C T  
 SEQ. ID. NO. 36 A G G A A A G C A A G G C T C T C T T T A G A A C  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 G A G A A G A A A A T G G A T A C A C T G C T T T  
 SEQ. ID. NO. 46 C T T T A G A A C A A T T A T C A C A T A C C C C  
 SEQ. ID. NO. 36 A A T T A T C A C A T A C C C C T G G T T C C A G  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 G A A A A T G T C A C C T C T A T C A T G T T T C  
 SEQ. ID. NO. 46 T G G T T C C A G A A C T C C T C G G T T A T T C  
 SEQ. ID. NO. 36 A A C T C C T C G G T T A T T C T G T T C T T A A  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 T A G T A G C G C T T A G T G A A T A T G A T C A  
 SEQ. ID. NO. 46 T G T T C T T A A A C A A G A A A G A T C T T C T  
 SEQ. ID. NO. 36 A C A A G A A A G A T C T T C T A G A G G A G A A  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 A G T T C T C G T G G A G T C A G A C A A T G A G  
 SEQ. ID. NO. 46 A G A G G A G A A A A T C A T G T A T T C C C A T  
 SEQ. ID. NO. 36 A A T C A T G T A T T C C C A T C T A G T C G A C  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 A A C C G A A T G G A G G A A A G C A A G G C T C  
 SEQ. ID. NO. 46 C T A G T C G A C T A C T T C C C A G A A T A T G  
 SEQ. ID. NO. 36 T A C T T C C C A G A A T A T G A T G G A C C C C  
 SEQ. ID. NO. 32

Figure 11t

SEQ. ID. NO. 40 T C T T T A G A A C A A T T A T C A C A T A C C C  
 SEQ. ID. NO. 46 A T G G A C C C C A G A G A G A T G C C C A G G C  
 SEQ. ID. NO. 36 A G A G A G A T G C C C A G G C A G C C C G A G A  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 C T G G T T C C A G A A C T C C T C G G T T A T T  
 SEQ. ID. NO. 46 A G C C C G A G A A T T C A T T C T G A A G A T G  
 SEQ. ID. NO. 36 A T T C A T T C T G A A G A T G T T C G T G G A C  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 C T G T T C T T A A A C A A G A A A G A T C T T C  
 SEQ. ID. NO. 46 T T C G T G G A C C T G A A C C C A G A C A G T G  
 SEQ. ID. NO. 36 C T G A A C C C A G A C A G T G A C A A A A T T A  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 T A G A G G A G A A A A T C A T G T A T T C C C A  
 SEQ. ID. NO. 46 A C A A A A T T A T C T A C T C C C A C T T C A C  
 SEQ. ID. NO. 36 T C T A C T C C C A C T T C A C G T G C G C C A C  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 T C T A G T C G A C T A C T T C C C A G A A T A T  
 SEQ. ID. NO. 46 G T G C G C C A C A G A C A C C G A G A A T A T C  
 SEQ. ID. NO. 36 A G A C A C C G A G A A T A T C C G C T T T G T C  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 G A T G G A C C C C A G A G A G A T G C C C A G G  
 SEQ. ID. NO. 46 C G C T T T G T C T T T G C T G C C G T C A A G G  
 SEQ. ID. NO. 36 T T T G C T G C C G T C A A G G A C A C C A T C C  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 C A G C C C G A G A A T T C A T T C T G A A G A T  
 SEQ. ID. NO. 46 A C A C C A T C C T C C A G T T G A A C C T G A A  
 SEQ. ID. NO. 36 T C C A G T T G A A C C T G A A G G A C T G C G G  
 SEQ. ID. NO. 32

SEQ. ID. NO. 40 G T T C G T G G A C C T G A A C C C A G A C A G T  
 SEQ. ID. NO. 46 G G A C T G C G G T C T G T T C T A A T T G T G C  
 SEQ. ID. NO. 36 T C T G T T C T A A  
 SEQ. ID. NO. 32

Figure 11u

SEQ. ID. NO. 40 G A C A A A A T T A T C T A C T C C C A C T T C A  
SEQ. ID. NO. 46 C T C C T A G A C A C C C G C C C T G C C C T T C  
SEQ. ID. NO. 36  
SEQ. ID. NO. 32

SEQ. ID. NO. 40 C G T G C G C C A C A G A C A C C G A G A A T A T  
SEQ. ID. NO. 46 C C T G G T  
SEQ. ID. NO. 36  
SEQ. ID. NO. 32

SEQ. ID. NO. 40 C C G C T T T G T C T T T G C T G C C G T C A A G  
SEQ. ID. NO. 46  
SEQ. ID. NO. 36  
SEQ. ID. NO. 32

SEQ. ID. NO. 40 G A C A C C A T C C T C C A G T T G A A C C T G A  
SEQ. ID. NO. 46  
SEQ. ID. NO. 36  
SEQ. ID. NO. 32

SEQ. ID. NO. 40 A G G A C T G C G G T C T G T T C T A A  
SEQ. ID. NO. 46  
SEQ. ID. NO. 36  
SEQ. ID. NO. 32

Figure 11v

# ClustalW Formatted Alignments

SEQ. ID. NO. 41 M V C E G K R S A S C P C F F L L T A K F Y W I L  
 SEQ. ID. NO. 47 M G S L L A L P A L L L L W G A V A E G P A K K V  
 SEQ. ID. NO. 37 M G S L L A L P A L L L L W G A V A E G P A K K V  
 SEQ. ID. NO. 33 M A F Y S C C W V L L A L T W H T S A Y G P D Q R

SEQ. ID. NO. 41 T M M Q R T H S Q E Y A H S I R V D G D I I L G G  
 SEQ. ID. NO. 47 L T L E G D L V L G G L F P V H Q K G G P A E D C  
 SEQ. ID. NO. 37 L T L E G D L V L G G L F P V H Q K G G P A E D C  
 SEQ. ID. NO. 33 A Q K K G D I I L G G L F P I H F G V A A K D Q D

SEQ. ID. NO. 41 L F P V H A K G E R G V P C G E L K K E K G I H R  
 SEQ. ID. NO. 47 G P V N E H R G I Q R L E A M L F A L D R I N R D  
 SEQ. ID. NO. 37 G P V N E H R G I Q R L E A M L F A L D R I N R D  
 SEQ. ID. NO. 33 L K S R P E S V E C I R Y N F R G F R W L Q A M I

SEQ. ID. NO. 41 L E A M L Y A I D Q I N K D P D L L S N I T L G V  
 SEQ. ID. NO. 47 P H L L P G V R L G A H I L D S C S K D T H A L E  
 SEQ. ID. NO. 37 P H L L P G V R L G A H I L D S C S K D T H A L E  
 SEQ. ID. NO. 33 F A I E E I N S S P A L L P N L T L G Y R I F D T

SEQ. ID. NO. 41 R I L D T C S R D T Y A L E Q S L T F V Q A L I E  
 SEQ. ID. NO. 47 Q A L D F V R A S L S R G A D G S R H I C P D G S  
 SEQ. ID. NO. 37 Q A L D F V R A S L S R G A D G S R H I C P D G S  
 SEQ. ID. NO. 33 C N T V S K A L E A T L S F V A Q N K I D S L N L

SEQ. ID. NO. 41 K D A S D V K C A N G D P P I F T K P D K I S G V  
 SEQ. ID. NO. 47 Y A T H G D A P T A I T G V I G G S Y S D V S I Q  
 SEQ. ID. NO. 37 Y A T H G D A P T A I T G V I G G S Y S D V S I Q  
 SEQ. ID. NO. 33 D E F C N C S E H I P S T I A V V G A T G S G V S

SEQ. ID. NO. 41 I G A A A S S V S I M V A N I L R L F K I P Q I S  
 SEQ. ID. NO. 47 V A N L L R L F Q I P Q I S Y A S T S A K L S D K  
 SEQ. ID. NO. 37 V A N L L R L F Q I P Q I S Y A S T S A K L S D K  
 SEQ. ID. NO. 33 T A V A N L L G L F Y I P Q V S Y A S S S R L L S

SEQ. ID. NO. 41 Y A S T A P E L S D N T R Y D F F S R V V P P D S  
 SEQ. ID. NO. 47 S R Y D Y F A R T V P P D F F Q A K A M A E I L R  
 SEQ. ID. NO. 37 S R Y D Y F A R T V P P D F F Q A K A M A E I L R  
 SEQ. ID. NO. 33 N K N Q F K S F L R T I P N D E H Q A T A M A D I

Figure 12a

SEQ. ID. NO. 41 YQAQAMVDIVTALGWNYVSTLASEG  
 SEQ. ID. NO. 47 FFNWTYVSTVASEGDYGETGIEAFE  
 SEQ. ID. NO. 37 FFNWTYVSTVASEGDYGETGIEAFE  
 SEQ. ID. NO. 33 IEYFRWNWVGTTAADDYGRPGIEK

SEQ. ID. NO. 41 NYGESGV E AFTQISREIGGVCI AQS  
 SEQ. ID. NO. 47 LEARARNICVATSEKVGGRAMSRAAF  
 SEQ. ID. NO. 37 LEARARNICVATSEKVGGRAMSRAAF  
 SEQ. ID. NO. 33 FREEAEERDICI D FSELISQYSDEE

SEQ. ID. NO. 41 QKIPREPRPGEF EK I I K R L L E T P N A  
 SEQ. ID. NO. 47 EG V V R A L L Q K P S A R V A V L F T R S E D A  
 SEQ. ID. NO. 37 EG V V R A L L Q K P S A R V A V L F T R S E D A  
 SEQ. ID. NO. 33 E I Q H V V E V I Q N S T A K V I V V F S S G P D

SEQ. ID. NO. 41 R A V I M F A N E D D I R R I L E A A K K L N Q S  
 SEQ. ID. NO. 47 R E L L A A S Q R L N A S F T W V A S D G W G A L  
 SEQ. ID. NO. 37 R E L L A A S Q R L N A S F T W V A S D G W G A L  
 SEQ. ID. NO. 33 L E P L I K E I V R R N I T G K I W L A S E A W A

SEQ. ID. NO. 41 G H F L W I G S D S W G S K I A P V Y Q Q E E I A  
 SEQ. ID. NO. 47 E S V V A G S E G A A E G A I T I E L A S Y P I S  
 SEQ. ID. NO. 37 E S V V A G S E G A A E G A I T I E L A S Y P I S  
 SEQ. ID. NO. 33 S S S L I A M P Q Y F H V V G G T I G F A L K A G

SEQ. ID. NO. 41 E G A V T I L P K R A S I D G F D R Y F R S R T L  
 SEQ. ID. NO. 47 D F A S Y F Q S L D P W N N S R N P W F R E F W E  
 SEQ. ID. NO. 37 D F A S Y F Q S L D P W N N S R N P W F R E F W E  
 SEQ. ID. NO. 33 Q I P G F R E F L K K V H P R K S V H N G F A K E

SEQ. ID. NO. 41 A N N R R N V W F A E F W E E N F G C K L G S H G  
 SEQ. ID. NO. 47 Q R F R C S F R Q R D C A A H S L R A V P F E Q E  
 SEQ. ID. NO. 37 Q R F R C S F R Q R D C A A H S L R A V P F E Q E  
 SEQ. ID. NO. 33 F W E E T F N C H L Q E G A K G P L P V D T F L R

SEQ. ID. NO. 41 K R N S H I K K C T G L E R I A R D S S Y E Q E G  
 SEQ. ID. NO. 47 S K I M F V V N A V Y A M A H A L H N M H R A L C  
 SEQ. ID. NO. 37 S K I M F V V N A V Y A M A H A L H N M H R A L C  
 SEQ. ID. NO. 33 G H E E S G D R F S N S S T A F R P L C T G D E N

Figure 12b

SEQ. ID. NO. 41 K V Q F V I D A V Y S M A Y A L H N M H K D L C P  
 SEQ. ID. NO. 47 P N T T R L C D A M R P V N G R R L Y K D F V L N  
 SEQ. ID. NO. 37 P N T T R L C D A M R P V N G R R L Y K D F V L N  
 SEQ. ID. NO. 33 I S S V E T P Y I D Y T H L R I S Y N V Y L A V Y

SEQ. ID. NO. 41 G Y I G L C P R M S T I D G K E L L G Y I R A V N  
 SEQ. ID. NO. 47 V K F D A P F R P A D T H N E V R F D R F G D G I  
 SEQ. ID. NO. 37 V K F D A P F R P A D T H N E V R F D R F G D G I  
 SEQ. ID. NO. 33 S I A H A L Q D I Y T C L P G R G L F T N G S C A

SEQ. ID. NO. 41 F N G S A G T P V T F N E N G D A P G R Y D I F Q  
 SEQ. ID. NO. 47 G R Y N I F T Y L R A G S G R Y R Y Q K V G Y W A  
 SEQ. ID. NO. 37 G R Y N I F T Y L R A G S G R Y R Y Q K V G Y W A  
 SEQ. ID. NO. 33 D I K K V E A W Q V L K H L R H L N F T N N M G E

SEQ. ID. NO. 41 Y Q I T N K S T E Y K V I G H W T N Q L H L K V E  
 SEQ. ID. NO. 47 E G L T L D T S L I P W A S P S A G P L P A S R C  
 SEQ. ID. NO. 37 E G L T L D T S L I P W A S P S A G P L P A S R C  
 SEQ. ID. NO. 33 Q V T F D E C G D L V G N Y S I I N W H L S P E D

SEQ. ID. NO. 41 D M Q W A H R E H T H P A S V C S L P C K P G E R  
 SEQ. ID. NO. 47 S E P C L Q N E V K S V Q P G E V C C W L C I P C  
 SEQ. ID. NO. 37 S E P C L Q N E V K S V Q P G E V C C W L C I P C  
 SEQ. ID. NO. 33 G S I V F K E V G Y Y N V Y A K K G E R L F I N E

SEQ. ID. NO. 41 K K T V K G V P C C W H C E R C E G Y N Y Q V D E  
 SEQ. ID. NO. 47 Q P Y E Y R L D E F T C A D C G L G Y W P N A S L  
 SEQ. ID. NO. 37 Q P Y E Y R L D E F T C A D C G L G Y W P N A S L  
 SEQ. ID. NO. 33 E K I L W S G F S R E V P F S N C S R D C L A G T

SEQ. ID. NO. 41 L S C E L C P L D Q R P N M N R T G C Q L I P I I  
 SEQ. ID. NO. 47 T G C F E L P Q E Y I R W G D A W A V G P V T I A  
 SEQ. ID. NO. 37 T G C F E L P Q E Y I R W G D A W A V G P V T I A  
 SEQ. ID. NO. 33 R K G I I E G E P T C C F E C V E C P D G E Y S D

SEQ. ID. NO. 41 K L E W H S P W A V V P V F V A I L G I I A T T F  
 SEQ. ID. NO. 47 C L G A L A T L F V L G V F V R H N A T P V V K A  
 SEQ. ID. NO. 37 C L G A L A T L F V L G V F V R H N A T P V V K A  
 SEQ. ID. NO. 33 E T D A S A C N K C P D D F W S N E N H T S C F E

Figure 12c



SEQ. ID. NO. 41 V I V T F V R Y N D T P I V R A S G R E L S Y V L  
 SEQ. ID. NO. 47 S G R E L C Y I L L G G V F L C Y C M T F I F I A  
 SEQ. ID. NO. 37 S G R E L C Y I L L G G V F L C Y C M T F I F I A  
 SEQ. ID. NO. 33 L P Q E Y I R W G D A W A V G P V T I A C L G A L

SEQ. ID. NO. 41 L T G I F L C Y S I T F L M I A A P D T I I C S F  
 SEQ. ID. NO. 47 K P S T A V C T L R R L G L G T A F S V C Y S A L  
 SEQ. ID. NO. 37 K P S T A V C T L R R L G L G T A F S V C Y S A L  
 SEQ. ID. NO. 33 A T L F V L G V F V R H N A T P V V K A S G R E L

SEQ. ID. NO. 41 R R V F L G L G M C F S Y A A L L T K T N R I H R  
 SEQ. ID. NO. 47 L T K T N R I A R I F G G A R E G A Q R P R F I S  
 SEQ. ID. NO. 37 L T K T N R I A R I F G G A R E G A Q R P R F I S  
 SEQ. ID. NO. 33 C Y I L L G G V F L C Y C M T F I F I A K P S T A

SEQ. ID. NO. 41 I F E Q G K K S V T A P K F I S P A S Q L V I T F  
 SEQ. ID. NO. 47 P A S Q V A I C L A L I S G Q L L I V V A W L V V  
 SEQ. ID. NO. 37 P A S Q V A I C L A L I S G Q L L I V V A W L V V  
 SEQ. ID. NO. 33 V C T L R R L G L G T A F S V C Y S A L L T K T N

SEQ. ID. NO. 41 S L I S V Q L L G V F V W F V V D P P H I I I D Y  
 SEQ. ID. NO. 47 E A P G T G K E T A P E R R E V V T L R C N H R D  
 SEQ. ID. NO. 37 E A P G T G K E T A P E R R E V V T L R C N H R D  
 SEQ. ID. NO. 33 R I A R I F G G A R E G A Q R P R F I S P A S Q V

SEQ. ID. NO. 41 G E Q R T L D P E K A R G V L K C D I S D L S L I  
 SEQ. ID. NO. 47 A S M L G S L A Y N V L L I A L C T L Y A F K T R  
 SEQ. ID. NO. 37 A S M L G S L A Y N V L L I A L C T L Y A F K T R  
 SEQ. ID. NO. 33 A I C L A L I S G Q L L I V V A W L V V E A P G T

SEQ. ID. NO. 41 C S L G Y S I L L M V T C T V Y A I K T R G V P E  
 SEQ. ID. NO. 47 K C P E N F N E A K F I G F T M Y T T C I I W L A  
 SEQ. ID. NO. 37 K C P E N F N E A K F I G F T M Y T T C I I W L A  
 SEQ. ID. NO. 33 G K E T A P E R R E V V T L R C N H R D A S M L G

SEQ. ID. NO. 41 T F N E A K P I G F T M Y T T C I I W L A F I P I  
 SEQ. ID. NO. 47 F L P I F Y V T S S D Y R V Q T T T M C V S V S L  
 SEQ. ID. NO. 37 F L P I F Y V T S S D Y R V Q T T T M C V S V S L  
 SEQ. ID. NO. 33 S L A Y N V L L I A L C T L Y A F N T R K C P E N

Figure 12d

SEQ. ID. NO. 41 F F G T A Q S A E K M Y I Q T T T L T V S M S L S  
 SEQ. ID. NO. 47 S G S V V L G C L F A P K L H I I L F Q P Q K N T  
 SEQ. ID. NO. 37 S G S V V L G C L F A P K L H I I L F Q P Q K N T  
 SEQ. ID. NO. 33 F N E A K F I G F T M Y T T C I I W L A L L P I F

SEQ. ID. NO. 41 A S V S L G M L Y M P K V Y I I I F H P E Q N T I  
 SEQ. ID. NO. 47 I E E V R C S T A A H A F K V A A R A T L R R S N  
 SEQ. ID. NO. 37 I E E V R C S T A A H A F K V A A R A T L R R S N  
 SEQ. ID. NO. 33 Y V T S S D Y R V Q T T T M C V S V S L S G S V V

SEQ. ID. NO. 41 E E V R C S T A A H A F K V A A R A T L R R S N V  
 SEQ. ID. NO. 47 V S R K R S S S L G G S T G S T P S S S I S S K S  
 SEQ. ID. NO. 37 V S R K R S S S L G G S T G S T P S S S I S S K S  
 SEQ. ID. NO. 33 L G C L F A P K L H I I L F Q P Q K N V V S H R A

SEQ. ID. NO. 41 S R K R S S S L G G S T G S T P S S S I S S K S N  
 SEQ. ID. NO. 47 N S E D P F P Q P E R Q K Q Q Q P L A L T Q Q E Q  
 SEQ. ID. NO. 37 N S E D P F P Q P E R Q K Q Q Q P L A L T Q Q E Q  
 SEQ. ID. NO. 33 P T S R F G S A A A R A S S S L G Q G S G S Q F V

SEQ. ID. NO. 41 S E D P F P Q P E R Q K Q Q Q P L A L T Q Q E Q Q  
 SEQ. ID. NO. 47 Q Q Q P L T L P Q Q Q R S Q Q Q P R C K Q K V I F  
 SEQ. ID. NO. 37 Q Q Q P L T L P Q Q Q R S Q Q Q P R C K Q K V I F  
 SEQ. ID. NO. 33 P T V C N G R E V V D S T T S S L M T L E S I M A

SEQ. ID. NO. 41 Q Q P L T L P Q Q Q R S Q Q Q P R C K Q K V I F G  
 SEQ. ID. NO. 47 G S G T V T F S L S F D E P Q K N A M A H G N S T  
 SEQ. ID. NO. 37 G S G T V T F S L S F D E P Q K N A M A H G N S T  
 SEQ. ID. NO. 33 C C L S E E A K E A R R I N D E I E R Q L R R D K

SEQ. ID. NO. 41 S G T V T F S L S F D E P Q K N A M A H G N S T H  
 SEQ. ID. NO. 47 H Q N S L E A Q K S S D T L T R H Q P L L P L Q C  
 SEQ. ID. NO. 37 H Q N S L E A Q K S S D T L T R H Q P L L P L Q C  
 SEQ. ID. NO. 33 R D A R R E L K L L L L G T G E S G K S T F I K Q

SEQ. ID. NO. 41 Q N S L E A Q K S S D T L T R H Q P L L P L Q C G  
 SEQ. ID. NO. 47 G E T D L D L T V Q E T G L Q G P V G G D Q R P E  
 SEQ. ID. NO. 37 G E T D L D L T V Q E T G L Q G P V G G D Q R P E  
 SEQ. ID. NO. 33 M R I I H G S G Y S D E D K R G F T K L V Y Q N I

Figure 12e

SEQ. ID. NO. 41 E T D L D L T V Q E T G L Q G P V G G D Q R P E V  
 SEQ. ID. NO. 47 V E D P E E L S P A L V V S S S Q S F V I S G G G  
 SEQ. ID. NO. 37 V E D P E E L S P A L V V S S S Q S F V I S G G G  
 SEQ. ID. NO. 33 F T A M Q A M I R A M D T L K I P Y K Y E H N K A

SEQ. ID. NO. 41 E D P E E L S P A L V V S S S Q S F V I S G G G S  
 SEQ. ID. NO. 47 S T V T E N V V N S A A A M T L E S I M A C C L S  
 SEQ. ID. NO. 37 S T V T E N V V N S M T L E S I M A C C L S E E A  
 SEQ. ID. NO. 33 H A Q L V R E V D V E K V S A F E N P Y V D A I K

SEQ. ID. NO. 41 T V T E N V V N S M T L E S I M A C C L S E E A K  
 SEQ. ID. NO. 47 E E A K E A R R I N D E I E R Q L R R D K R D A R  
 SEQ. ID. NO. 37 K E A R R I N D E I E R Q L R R D K R D A R R E L  
 SEQ. ID. NO. 33 S L W N D P G I Q E C Y D R R R E Y Q L S D S T K

SEQ. ID. NO. 41 E A R R I N D E I E R Q L R R D K R D A R R E L K  
 SEQ. ID. NO. 47 R E L K L L L L G T G E S G K S T F I K Q M R I I  
 SEQ. ID. NO. 37 K L L L L G T G E S G K S T F I K Q M R I I H G S  
 SEQ. ID. NO. 33 Y Y L N D L D R V A D P A Y L P T Q Q D V L R V R

SEQ. ID. NO. 41 L L L L G T G E S G K S T F I K Q M R I I H G S G  
 SEQ. ID. NO. 47 H G S G Y S D E D K R G F T K L V Y Q N I F T A M  
 SEQ. ID. NO. 37 G Y S D E D K R G F T K L V Y Q N I F T A M Q A M  
 SEQ. ID. NO. 33 V P T T G I I E Y P F D L Q S V I F R M V D V G G

SEQ. ID. NO. 41 Y S D E D K R G F T K L V Y Q N I F T A M Q A M I  
 SEQ. ID. NO. 47 Q A M I R A M D T L K I P Y K Y E H N K A H A Q L  
 SEQ. ID. NO. 37 I R A M D T L K I P Y K Y E H N K A H A Q L V R E  
 SEQ. ID. NO. 33 Q R S E R R K W I H C F E N V T S I M F L V A L S

SEQ. ID. NO. 41 R A M D T L K I P Y K Y E H N K A H A Q L V R E V  
 SEQ. ID. NO. 47 V R E V D V E K V S A F E N P Y V D A I K S L W N  
 SEQ. ID. NO. 37 V D V E K V S A F E N P Y V D A I K S L W N D P G  
 SEQ. ID. NO. 33 E Y D Q V L V E S D N E N R M E E S K A L F R T I

SEQ. ID. NO. 41 D V E K V S A F E N P Y V D A I K S L W N D P G I  
 SEQ. ID. NO. 47 D P G I Q E C Y D R R R E Y Q L S D S T K Y Y L N  
 SEQ. ID. NO. 37 I Q E C Y D R R R E Y Q L S D S T K Y Y L N D L D  
 SEQ. ID. NO. 33 I T Y P W F Q N S S V I L F L N K K D L L E E K I

Figure 12f

SEQ. ID. NO. 41 Q E C Y D R R R E Y Q L S D S T K Y Y L N D L D R  
 SEQ. ID. NO. 47 D L D R V A D P A Y L P T Q Q D V L R V R V P T T  
 SEQ. ID. NO. 37 R V A D P A Y L P T Q Q D V L R V R V P T T G I I  
 SEQ. ID. NO. 33 M Y S H L V D Y F P E Y D G P Q R D A Q A A R E F

SEQ. ID. NO. 41 V A D P A Y L P T Q Q D V L R V R V P T T G I I E  
 SEQ. ID. NO. 47 G I I E Y P F D L Q S V I F R M V D V G G Q R S E  
 SEQ. ID. NO. 37 E Y P F D L Q S V I F R M V D V G G Q R S E R R K  
 SEQ. ID. NO. 33 I L K M F V D L N P D S D K I I Y S H F T C A T D

SEQ. ID. NO. 41 Y P F D L Q S V I F R M V D V G G Q R S E R R K W  
 SEQ. ID. NO. 47 R R K W I H C F E N V T S I M F L V A L S E Y D Q  
 SEQ. ID. NO. 37 W I H C F E N V T S I M F L V A L S E Y D Q V L V  
 SEQ. ID. NO. 33 T E N I R F V F A A V K D T I L Q L N L K D C G L

SEQ. ID. NO. 41 I H C F E N V T S I M F L V A L S E Y D Q V L V E  
 SEQ. ID. NO. 47 V L V E S D N E N R M E E S K A L F R T I I T Y P  
 SEQ. ID. NO. 37 E S D N E N R M E E S K A L F R T I I T Y P W F Q  
 SEQ. ID. NO. 33 F

SEQ. ID. NO. 41 S D N E N R M E E S K A L F R T I I T Y P W F Q N  
 SEQ. ID. NO. 47 W F Q N S S V I L F L N K K D L L E E K I M Y S H  
 SEQ. ID. NO. 37 N S S V I L F L N K K D L L E E K I M Y S H L V D  
 SEQ. ID. NO. 33

SEQ. ID. NO. 41 S S V I L F L N K K D L L E E K I M Y S H L V D Y  
 SEQ. ID. NO. 47 L V D Y F P E Y D G P Q R D A Q A A R E F I L K M  
 SEQ. ID. NO. 37 Y F P E Y D G P Q R D A Q A A R E F I L K M F V D  
 SEQ. ID. NO. 33

SEQ. ID. NO. 41 F P E Y D G P Q R D A Q A A R E F I L K M F V D L  
 SEQ. ID. NO. 47 F V D L N P D S D K I I Y S H F T C A T D T E N I  
 SEQ. ID. NO. 37 L N P D S D K I I Y S H F T C A T D T E N I R F V  
 SEQ. ID. NO. 33

SEQ. ID. NO. 41 N P D S D K I I Y S H F T C A T D T E N I R F V F  
 SEQ. ID. NO. 47 R F V F A A V K D T I L Q L N L K D C G L F  
 SEQ. ID. NO. 37 F A A V K D T I L Q L N L K D C G L F  
 SEQ. ID. NO. 33

Figure 12g

SEQ. ID. NO. 41 A A V K D T I L Q L N L K D C G L F  
SEQ. ID. NO. 47  
SEQ. ID. NO. 37  
SEQ. ID. NO. 33

Figure 12h

### ClustalW Formatted Alignments

SEQ. ID. NO. 44 A T G T T G C T G C T G C T A C T G G C G C  
SEQ. ID. NO. 42 A T G G C T T C C C C G C G G A G C T C C G G G C

SEQ. ID. NO. 44 C A C T C T T C C T C C G C C C C C G G G C G C  
SEQ. ID. NO. 42 A G C C C G G G C C G C C G C C G C C G C C G C C

SEQ. ID. NO. 44 G G G C G G G G C G C A G A C C C C C A A C G C C  
SEQ. ID. NO. 42 A C C G C C G C C C G C G C G C C T G C T A C T G

SEQ. ID. NO. 44 A C C T C A G A A G G T T G C C A G A T C A T A C  
SEQ. ID. NO. 42 C T A C T G C T G C T G C C G C T G C T G C T G C

SEQ. ID. NO. 44 A C C C G C C C T G G G A A G G G G G C A T C A G  
SEQ. ID. NO. 42 C T C T G G C G C C C G G G G C C T G G G G C T G

SEQ. ID. NO. 44 G T A C C G G G G C C T G A C T C G G G A C C A G  
SEQ. ID. NO. 42 G G C G C G G G G C G C C C C C C G G C C G C C G

SEQ. ID. NO. 44 G T G A A G G C T A T C A A C T T C C T G C C A G  
SEQ. ID. NO. 42 C C C A G C A G C C C G C C G C T C T C C A T C A

SEQ. ID. NO. 44 T G G A C T A T G A G A T T G A G T A T G T G T G  
SEQ. ID. NO. 42 T G G G C C T C A T G C C G C T C A C C A A G G A

SEQ. ID. NO. 44 C C G G G G G G A G C G C G A G G T G G T G G G G  
SEQ. ID. NO. 42 G G T G G C C A A G G G C A G C A T C G G G C G C

SEQ. ID. NO. 44 C C C A A G G T C C G C A A G T G C C T G G C C A  
SEQ. ID. NO. 42 G G T G T G C T C C C C G C C G T G G A A C T G G

SEQ. ID. NO. 44 A C G G C T C C T G G A C A G A T A T G G A C A C  
SEQ. ID. NO. 42 C C A T C G A G C A G A T C C G C A A C G A G T C

SEQ. ID. NO. 44 A C C C A G C C G C T G T G T C C G A A T C T G C  
SEQ. ID. NO. 42 A C T C C T G C G C C C T T A C T T C C T C G A C

Figure 13a

SEQ. ID. NO. 44 T C C A A G T C T T A T T T G A C C C T G G A A A  
SEQ. ID. NO. 42 C T G C G G C T C T A T G A C A C G G A G T G C G

SEQ. ID. NO. 44 A T G G G A A G G T T T C C T G A C G G G T G G  
SEQ. ID. NO. 42 A C A A C G C A A A A G G G T T G A A A G C C T T

SEQ. ID. NO. 44 G G A C C T C C C A G C T C T G G A C G G A G C C  
SEQ. ID. NO. 42 C T A C G A T G C A A T A A A A T A C G G G C C G

SEQ. ID. NO. 44 C G G G T G G A T T T C C G G T G T G A C C C C G  
SEQ. ID. NO. 42 A A C C A C T T G A T G G T G T T T G G A G G C G

SEQ. ID. NO. 44 A C T T C C A T C T G G T G G G C A G C T C C C G  
SEQ. ID. NO. 42 T C T G T C C A T C C G T C A C A T C C A T C A T

SEQ. ID. NO. 44 G A G C A T C T G T A G T C A G G G C C A G T G G  
SEQ. ID. NO. 42 T G C A G A G T C C C T C C A A G G C T G G A A T

SEQ. ID. NO. 44 A G C A C C C C C A A G C C C C A C T G C C A G G  
SEQ. ID. NO. 42 C T G G T G C A G C T T T C T T T T G C T G C A A

SEQ. ID. NO. 44 T G A A T C G A A C G C C A C A C T C A G A A C G  
SEQ. ID. NO. 42 C C A C G C C T G T T C T A G C C G A T A A G A A

SEQ. ID. NO. 44 G C G C G C A G T G T A C A T C G G G G C A C T G  
SEQ. ID. NO. 42 A A A A T A C C C T T A T T T C T T T C G G A C C

SEQ. ID. NO. 44 T T T C C C A T G A G C G G G G G C T G G C C A G  
SEQ. ID. NO. 42 G T C C C A T C A G A C A A T G C G G T G A A T C

SEQ. ID. NO. 44 G G G G C C A G G C C T G C C A G C C C G C G G T  
SEQ. ID. NO. 42 C A G C C A T T C T G A A G T T G C T C A A G C A

SEQ. ID. NO. 44 G G A G A T G G C G C T G G A G G A C G T G A A T  
SEQ. ID. NO. 42 C T A C C A G T G G A A G C G C G T G G G C A C G

SEQ. ID. NO. 44 A G C C G C A G G G A C A T C C T G C C G G A C T  
SEQ. ID. NO. 42 C T G A C G C A A G A C G T T C A G A G G T T C T

Figure 13b

SEQ. ID. NO. 44 A T G A G C T C A A G C T C A T C C A C C A C G A  
SEQ. ID. NO. 42 C T G A G G T G C G G A A T G A C C T G A C T G G

SEQ. ID. NO. 44 C A G C A A G T G T G A T C C A G G C C A A G C C  
SEQ. ID. NO. 42 A G T T C T G T A T G G C G A G G A C A T T G A G

SEQ. ID. NO. 44 A C C A A G T A C C T A T A T G A G C T G C T C T  
SEQ. ID. NO. 42 A T T T C A G A C A C C G A G A G C T T C T C C A

SEQ. ID. NO. 44 A C A A C G A C C C T A T C A A G A T C A T C C T  
SEQ. ID. NO. 42 A C G A T C C C T G T A C C A G T G T C A A A A A

SEQ. ID. NO. 44 T A T G C C T G G C T G C A G C T C T G T C T C C  
SEQ. ID. NO. 42 G C T G A A G G G G A A T G A T G T G C G G A T C

SEQ. ID. NO. 44 A C G C T G G T G G C T G A G G C T G C T A G G A  
SEQ. ID. NO. 42 A T C C T T G G C C A G T T T G A C C A G A A T A

SEQ. ID. NO. 44 T G T G G A A C C T C A T T G T G C T T T C C T A  
SEQ. ID. NO. 42 T G G C A G C A A A A G T G T T C T G T T G T G C

SEQ. ID. NO. 44 T G G C T C C A G C T C A C C A G C C C T G T C A  
SEQ. ID. NO. 42 A T A C G A G G A G A A C A T G T A T G G T A G T

SEQ. ID. NO. 44 A A C C G G C A G C G T T T C C C C A C T T T C T  
SEQ. ID. NO. 42 A A A T A T C A G T G G A T C A T T C C G G G C T

SEQ. ID. NO. 44 T C C G A A C G C A C C C A T C A G C C A C A C T  
SEQ. ID. NO. 42 G G T A C G A G C C T T C T T G G T G G G A G C A

SEQ. ID. NO. 44 C C A C A A C C C T A C C C G C G T G A A A C T C  
SEQ. ID. NO. 42 G G T G C A C A C G G A A G C C A A C T C A T C C

SEQ. ID. NO. 44 T T T G A A A A G T G G G G C T G G A A G A A G A  
SEQ. ID. NO. 42 C G C T G C C T C C G G A A G A A T C T G C T T G

SEQ. ID. NO. 44 T T G C T A C C A T C C A G C A G A C C A C T G A  
SEQ. ID. NO. 42 C T G C C A T G G A G G G C T A C A T T G G C G T

Figure 13c



SEQ. ID. NO. 44 GGTCTTCTCACTTCGACTCTGGACGAC  
SEQ. ID. NO. 42 GGATTTTCGAGCCCCCTGAGCTCCAAG

SEQ. ID. NO. 44 CTGGAGGAACGAGTGAAGGAGGCTG  
SEQ. ID. NO. 42 CAGATCAAGACCATCTCAGGAAAGA

SEQ. ID. NO. 44 GAATTGAGATTACTTTCCGCCAGAG  
SEQ. ID. NO. 42 CTCCACAGCAGTATGAGAGAGAGTA

SEQ. ID. NO. 44 TTTCTTCTCAGATCCAGCTGTGCCC  
SEQ. ID. NO. 42 CAACAACAAGCGGTCAGGCGTGGGG

SEQ. ID. NO. 44 GTCAAAAACCTGAAGCGCCAGGATG  
SEQ. ID. NO. 42 CCCAGCAAGTTCCACGGGTACGCCCT

SEQ. ID. NO. 44 CCCGAATCATCTGTTGGGACTTTTCTA  
SEQ. ID. NO. 42 ACGATGGCATCTGGGGTCATCGCCAA

SEQ. ID. NO. 44 TGAGACTGAAGCCCCGGAAAGTTTTT  
SEQ. ID. NO. 42 GACACTGCAGAGGGCCATGGAGACA

SEQ. ID. NO. 44 TGTGAGGTGTACAAGGAGCGTCTCT  
SEQ. ID. NO. 42 CTGCATGCCAGCAGCCGGGCACCAAGC

SEQ. ID. NO. 44 TTGGGAAGAAGTACGTCTGGTTTCCT  
SEQ. ID. NO. 42 GGATCCAGGACTTCAACTACACGGA

SEQ. ID. NO. 44 CATTTGGGTGGTATGCTGACAAATTGG  
SEQ. ID. NO. 42 CCACACGCTGGGGCAGGATCATCCTC

SEQ. ID. NO. 44 TTCAAGATCTACGACCCTTCTATCA  
SEQ. ID. NO. 42 AATGCCATGAACGAGACCAACTTCT

SEQ. ID. NO. 44 ACTGCACAGTGGATGAGATGACTGA  
SEQ. ID. NO. 42 TCGGGGTCAACGGGTCAAGTTGTATT

SEQ. ID. NO. 44 GGCGGTGGAGGGCCACATCACAACT  
SEQ. ID. NO. 42 CCGGAATGGGGAGAGAAATGGGGACC

Figure 13d

SEQ. ID. NO. 44 G A G A T T G T C A T G C T G A A T C C T G C C A  
SEQ. ID. NO. 42 A T T A A A T T T A C T C A A T T T C A A G A C A

SEQ. ID. NO. 44 A T A C C C G C A G C A T T T C C A A C A T G A C  
SEQ. ID. NO. 42 G C A G G G A G G T G A A G G T G G G A G A G T A

SEQ. ID. NO. 44 A T C C C A G G A A T T T G T G G A G A A A C T A  
SEQ. ID. NO. 42 C A A C G C T G T G G C C G A C A C A C T G G A G

SEQ. ID. NO. 44 A C C A A G C G A C T G A A A A G A C A C C C T G  
SEQ. ID. NO. 42 A T C A T C A A T G A C A C C A T C A G G T T C C

SEQ. ID. NO. 44 A G G A G A C A G G A G G C T T C C A G G A G G C  
SEQ. ID. NO. 42 A A G G A T C C G A A C C A C C A A A A G A C A A

SEQ. ID. NO. 44 A C C G C T G G C C T A T G A T G C C A T C T G G  
SEQ. ID. NO. 42 G A C C A T C A T C C T G G A G C A G C T G C G G

SEQ. ID. NO. 44 G C C T T G G C A C T G G C C C T G A A C A A G A  
SEQ. ID. NO. 42 A A G A T C T C C C T A C C T C T C T A C A G C A

SEQ. ID. NO. 44 C A T C T G G A G G A G G C G G C C G T T C T G G  
SEQ. ID. NO. 42 T C C T C T C T G C C C T C A C C A T C C T C G G

SEQ. ID. NO. 44 T G T G C G C C T G G A G G A C T T C A A C T A C  
SEQ. ID. NO. 42 G A T G A T C A T G G C C A G T G C T T T T C T C

SEQ. ID. NO. 44 A A C A A C C A G A C C A T T A C C G A C C A A A  
SEQ. ID. NO. 42 T T C T T C A A C A T C A A G A A C C G G A A T C

SEQ. ID. NO. 44 T C T A C C G G G C A A T G A A C T C T T C G T C  
SEQ. ID. NO. 42 A G A A G C T C A T A A A G A T G T C G A G T C C

SEQ. ID. NO. 44 C T T T G A G G G T G T C T C T G G C C A T G T G  
SEQ. ID. NO. 42 A T A C A T G A A C A A C C T T A T C A T C C T T

SEQ. ID. NO. 44 G T G T T T G A T G C C A G C G G C T C T C G G A  
SEQ. ID. NO. 42 G G A G G G A T G C T C T C C T A T G C T T C C A

Figure 13e

SEQ. ID. NO. 44 TGGCATGGACGCTTATCGAGCAGCT  
SEQ. ID. NO. 42 TATTTCTCTTTTGGCCTTGATGGATC

SEQ. ID. NO. 44 TCAGGGTGGCAGCTACAAGAAGATT  
SEQ. ID. NO. 42 CTTTGTCTCTGTGAAAAGACCTTTTGA

SEQ. ID. NO. 44 GGCTACTATGACAGCACCAAGGATG  
SEQ. ID. NO. 42 ACACTTTGCAACCGTCAGGACCTTGA

SEQ. ID. NO. 44 ATCTTTCCTGGTCCAAAACAGATAA  
SEQ. ID. NO. 42 TTCTCACCGTGGGCTACACGACCGC

SEQ. ID. NO. 44 ATGGATTGGAGGGTCCCCCCCAGCT  
SEQ. ID. NO. 42 TTTTGGGGGCCATGTTTGTCAAAGACC

SEQ. ID. NO. 44 GACCAGACCCTGGTTCATCAAGACAT  
SEQ. ID. NO. 42 TGGAGAGTCCACGCCATCTTCAAAA

SEQ. ID. NO. 44 TCCGCTTTCCTGTTCACAGAAACTCTT  
SEQ. ID. NO. 42 ATGTGAAAATGAAGAAGAAGATCAT

SEQ. ID. NO. 44 TATCTCCGTCTCAGTTCTCTCCAGC  
SEQ. ID. NO. 42 CAAGGACCAGAAACTGCTTGTGATC

SEQ. ID. NO. 44 CTGGGCATTGTCCTAGCTGTTGTCT  
SEQ. ID. NO. 42 GTGGGGGGGCATGCTGCTGATCGACC

SEQ. ID. NO. 44 GTCTGTCCTTTAACATCTACAACCTC  
SEQ. ID. NO. 42 TGTGTATCCTGATCTGCTGGCAGGC

SEQ. ID. NO. 44 ACATGTCCGTTATATCCAGAACTCA  
SEQ. ID. NO. 42 TGTGGACCCCCCTGCGAAGGACAGTG

SEQ. ID. NO. 44 CAGCCCAACCTGAACAACCTGACTG  
SEQ. ID. NO. 42 GAGAAGTACAGCATGGAGCCGGACC

SEQ. ID. NO. 44 CTGTGGGGCTGCTCACTGGCTTTAGC  
SEQ. ID. NO. 42 CAGCAGGACGGGATATCTCCATCCG

Figure 13f

SEQ. ID. NO. 44 TGCTGTCTTCCCCCTGGGGCTCGAT  
SEQ. ID. NO. 42 CCTCTCTCTGGAGCACTGTGAGAAC

SEQ. ID. NO. 44 GGTTACCAACATTGGGAGGAACCACT  
SEQ. ID. NO. 42 ACCCATATGACCATCTGGCTTGGCA

SEQ. ID. NO. 44 TTCCTTTCGTCTGCCAGGCCCGCCT  
SEQ. ID. NO. 42 TCGTCTATGCCTACAAGGGACTTCT

SEQ. ID. NO. 44 CTGGCTCCTGGGCCTGGGCTTTAGT  
SEQ. ID. NO. 42 CATGTTGTTTCGGTTGTTTCTTAGCT

SEQ. ID. NO. 44 CTGGGCTACGGTTCCATGTTTCAACCA  
SEQ. ID. NO. 42 TGGGAGACCCGCAACGTCAGCATCC

SEQ. ID. NO. 44 AGATTTGGTGGGTCCACACGGTCTT  
SEQ. ID. NO. 42 CCGCACTCAACGACAGCAAGTACAT

SEQ. ID. NO. 44 CACAAAGAAGGAAGAAAAGAAGGAG  
SEQ. ID. NO. 42 CGGGATGAGTGCTACAACGTGGGG

SEQ. ID. NO. 44 TGGAGGAAGACTCTGGAACCCCTGGA  
SEQ. ID. NO. 42 ATCATGTGCATCATCGGGGGCCGCTG

SEQ. ID. NO. 44 AGCTGTATGCCACAGTGGGGCCTGCT  
SEQ. ID. NO. 42 TCTCCTTCTCTGACCCCGGGACCAAGCC

SEQ. ID. NO. 44 GGTGGGCATGGATGTCCTCACTCTC  
SEQ. ID. NO. 42 CAATGTGCAGTTCTGCATCGTGGCT

SEQ. ID. NO. 44 GCCATCTGGCAGATCGTGGACCCCTC  
SEQ. ID. NO. 42 CTGGTCATCATCTTCTGCAGCACCA

SEQ. ID. NO. 44 TGCACCGGACCAATTGAGACATTTGC  
SEQ. ID. NO. 42 TCACCCCTCTGCCTGGTATTCTGTGCC

SEQ. ID. NO. 44 CAAGGAGGAACCTAAGGAAGATATT  
SEQ. ID. NO. 42 GAAGCTCATCACCCCTGAGAACAAAC

Figure 13g

SEQ. ID. NO. 44 G A C G T C T C T A T T C T G C C C C A G C T G G  
SEQ. ID. NO. 42 C C A G A T G C A G C A A C G C A G A A C A G G C

SEQ. ID. NO. 44 A G C A T T G C A G C T C C A G G A A G A T G A A  
SEQ. ID. NO. 42 G A T T C C A G T T C A C T C A G A A T C A G A A

SEQ. ID. NO. 44 T A C A T G G C T T G G C A T T T T C T A T G G T  
SEQ. ID. NO. 42 G A A A G A A G A T T C T A A A A C G T C C A C C

SEQ. ID. NO. 44 T A C A A G G G G C T G C T G C T G C T G C T G G  
SEQ. ID. NO. 42 T C G G T C A C C A G T G T G A A C C A A G C C A

SEQ. ID. NO. 44 G A A T C T T C C T T G C T T A T G A G A C C A A  
SEQ. ID. NO. 42 G C A C A T C C C G C C T G G A G G G C C T A C A

SEQ. ID. NO. 44 G A G T G T G T C C A C T G A G A A G A T C A A T  
SEQ. ID. NO. 42 G T C A G A A A A C C A T C G C C T G C G A A T G

SEQ. ID. NO. 44 G A T C A C C G G G C T G T G G G C A T G G C T A  
SEQ. ID. NO. 42 A A G A T C A C A G A G C T G G A T A A A G A C T

SEQ. ID. NO. 44 T C T A C A A T G T G G C A G T C C T G T G C C T  
SEQ. ID. NO. 42 T G G A A G A G G T C A C C A T G C A G C T G C A

SEQ. ID. NO. 44 C A T C A C T G C T C C T G T C A C C A T G A T T  
SEQ. ID. NO. 42 G G A C A C A C C A G A A A A G A C C A C C T A C

SEQ. ID. NO. 44 C T G T C C A G C C A G C A G G A T G C A G C C T  
SEQ. ID. NO. 42 A T T A A A C A G A A C C A C T A C C A A G A G C

SEQ. ID. NO. 44 T T G C C T T T G C C T C T C T T G C C A T A G T  
SEQ. ID. NO. 42 T C A A T G A C A T C C T C A A C C T G G G A A A

SEQ. ID. NO. 44 T T T C T C C T C C T A T A T C A C T C T T G T T  
SEQ. ID. NO. 42 C T T C A C T G A G A G C A C A G A T G G A G G A

SEQ. ID. NO. 44 G T G C T C T T T G T G C C C A A G A T G C G C A  
SEQ. ID. NO. 42 A A G G C C A T T T T A A A A A A T C A C C T C G

Figure 13h

SEQ. ID. NO. 44 G G C T G A T C A C C C G A G G G G A A T G G C A  
 SEQ. ID. NO. 42 A T C A A A A T C C C C A G C T A C A G T G G A A

SEQ. ID. NO. 44 G T C G G A G G C G C A G G A C A C C A T G A A G  
 SEQ. ID. NO. 42 C A C A A C A G A G C C C T C T C G A A C A T G C

SEQ. ID. NO. 44 A C A G G G T C A T C G A C C A A C A A C A A C G  
 SEQ. ID. NO. 42 A A A G A T C C T A T A G A A G A T A T A A A C T

SEQ. ID. NO. 44 A G G A G G A G A A G T C C C G G C T G T T G G A  
 SEQ. ID. NO. 42 C T C C A G A A C A C A T C C A G C G T C G G C T

SEQ. ID. NO. 44 G A A G G A G A A C C G T G A A C T G G A A A A G  
 SEQ. ID. NO. 42 G T C C C T C C A G C T C C C C A T C C T C C A C

SEQ. ID. NO. 44 A T C A T T G C T G A G A A A G A G G A G C G T G  
 SEQ. ID. NO. 42 C A C G C C T A C C T C C C A T C C A T C G G A G

SEQ. ID. NO. 44 T C T C T G A A C T G C G C C A T C A A C T C C A  
 SEQ. ID. NO. 42 G C G T G G A C G C C A G C T G T G T C A G C C C

SEQ. ID. NO. 44 G T C T C G G C A G C A G C T C C G C T C C C G G  
 SEQ. ID. NO. 42 C T G C G T C A G C C C C A C C G C C A G C C C C

SEQ. ID. NO. 44 C G C C A C C C A C C G A C A C C C C C A G A A C  
 SEQ. ID. NO. 42 C G C C A C A G A C A T G T G C C A C C C T C C T

SEQ. ID. NO. 44 C C T C T G G G G G C C T G C C C A G G G G A C C  
 SEQ. ID. NO. 42 T C C G A G T C A T G G T C T C G G G C C T G G C

SEQ. ID. NO. 44 C C C T G A G C C C C C G A C C G G C T T A G C  
 SEQ. ID. NO. 42 G G C C G C C A T G A C T C T G G A G T C C A T C

SEQ. ID. NO. 44 T G T G A T G G G A G T C G A G T G C A T T T G C  
 SEQ. ID. NO. 42 A T G G C G T G C T G C C T G A G C G A G G A G G

SEQ. ID. NO. 44 T T T A T A A G G C G G C C G C C A T G A C T C T  
 SEQ. ID. NO. 42 C C A A G G A A G C C C G G C G G A T C A A C G A

Figure 13i

SEQ. ID. NO. 44 G G A G T C C A T C A T G G C G T G C T G C C T G  
 SEQ. ID. NO. 42 C G A G A T C G A G C G G C A G C T C C G C A G G

SEQ. ID. NO. 44 A G C G A G G A G G C C A A G G A A G C C C G G C  
 SEQ. ID. NO. 42 G A C A A G C G G G A C G C C C G C C G G G A G C

SEQ. ID. NO. 44 G G A T C A A C G A C G A G A T C G A G C G G C A  
 SEQ. ID. NO. 42 T C A A G C T G C T G C T G C T C G G G A C A G G

SEQ. ID. NO. 44 G C T C C G C A G G G A C A A G C G G G A C G C C  
 SEQ. ID. NO. 42 A G A G A G T G G C A A G A G T A C G T T T A T C

SEQ. ID. NO. 44 C G C C G G G A G C T C A A G C T G C T G C T G C  
 SEQ. ID. NO. 42 A A G C A G A T G A G A A T C A T C C A T G G G T

SEQ. ID. NO. 44 T C G G G A C A G G A G A G A G T G G C A A G A G  
 SEQ. ID. NO. 42 C A G G A T A C T C T G A T G A A G A T A A A A G

SEQ. ID. NO. 44 T A C G T T T A T C A A G C A G A T G A G A A T C  
 SEQ. ID. NO. 42 G G G C T T C A C C A A G C T G G T G T A T C A G

SEQ. ID. NO. 44 A T C C A T G G G T C A G G A T A C T C T G A T G  
 SEQ. ID. NO. 42 A A C A T C T T C A C G G C C A T G C A G G C C A

SEQ. ID. NO. 44 A A G A T A A A A G G G G C T T C A C C A A G C T  
 SEQ. ID. NO. 42 T G A T C A G A G C C A T G G A C A C A C T C A A

SEQ. ID. NO. 44 G G T G T A T C A G A A C A T C T T C A C G G C C  
 SEQ. ID. NO. 42 G A T C C C A T A C A A G T A T G A G C A C A A T

SEQ. ID. NO. 44 A T G C A G G C C A T G A T C A G A G C C A T G G  
 SEQ. ID. NO. 42 A A G G C T C A T G C A C A A T T A G T T C G A G

SEQ. ID. NO. 44 A C A C A C T C A A G A T C C C A T A C A A G T A  
 SEQ. ID. NO. 42 A A G T T G A T G T G G A G A A G G T G T C T G C

SEQ. ID. NO. 44 T G A G C A C A A T A A G G C T C A T G C A C A A  
 SEQ. ID. NO. 42 T T T T G A G A A T C C A T A T G T A G A T G C A

Figure 13j

SEQ. ID. NO. 44 T T A G T T C G A G A A G T T G A T G T G G A G A  
SEQ. ID. NO. 42 A T A A A G A G T T T A T G G A A T G A T C C T G

SEQ. ID. NO. 44 A G G T G T C T G C T T T T G A G A A T C C A T A  
SEQ. ID. NO. 42 G A A T C C A G G A A T G C T A T G A T A G A C G

SEQ. ID. NO. 44 T G T A G A T G C A A T A A A G A G T T T A T G G  
SEQ. ID. NO. 42 A C G A G A A T A T C A A T T A T C T G A C T C T

SEQ. ID. NO. 44 A A T G A T C C T G G A A T C C A G G A A T G C T  
SEQ. ID. NO. 42 A C C A A A T A C T A T C T T A A T G A C T T G G

SEQ. ID. NO. 44 A T G A T A G A C G A C G A G A A T A T C A A T T  
SEQ. ID. NO. 42 A C C G C G T A G C T G A C C C T G C C T A C C T

SEQ. ID. NO. 44 A T C T G A C T C T A C C A A A T A C T A T C T T  
SEQ. ID. NO. 42 G C C T A C G C A A C A A G A T G T G C T T A G A

SEQ. ID. NO. 44 A A T G A C T T G G A C C G C G T A G C T G A C C  
SEQ. ID. NO. 42 G T T C G A G T C C C C A C C A C A G G G A T C A

SEQ. ID. NO. 44 C T G C C T A C C T G C C T A C G C A A C A A G A  
SEQ. ID. NO. 42 T C G A A T A C C C C T T T G A C T T A C A A A G

SEQ. ID. NO. 44 T G T G C T T A G A G T T C G A G T C C C C A C C  
SEQ. ID. NO. 42 T G T C A T T T T T C A G A A T G G T C G A T G T A

SEQ. ID. NO. 44 A C A G G G A T C A T C G A A T A C C C C T T T G  
SEQ. ID. NO. 42 G G G G G C C A A A G G T C A G A G A G A A G A A

SEQ. ID. NO. 44 A C T T A C A A A G T G T C A T T T T T C A G A A T  
SEQ. ID. NO. 42 A A T G G A T A C A C T G C T T T T G A A A A T G T

SEQ. ID. NO. 44 G G T C G A T G T A G G G G G C C A A A G G T C A  
SEQ. ID. NO. 42 C A C C T C T A T C A T G T T T C T A G T A G C G

SEQ. ID. NO. 44 G A G A G A A G A A A A T G G A T A C A C T G C T  
SEQ. ID. NO. 42 C T T A G T G A A T A T G A T C A A G T T C T G G

Figure 13k



SEQ. ID. NO. 44 TTGAAAATGTCACCTCTATCATGTT  
SEQ. ID. NO. 42 TGGAGTCAGACAAATGAGAACCGAAAT

SEQ. ID. NO. 44 TCTAGTAGCGCTTAGTGAAATATGAT  
SEQ. ID. NO. 42 GGAGGAAAGCAAGGCTCTCTTTAGA

SEQ. ID. NO. 44 CAAGTTCTCTCGTGGAGTCAGACAAATG  
SEQ. ID. NO. 42 ACAATTATCACATACCCCTGGTTCC

SEQ. ID. NO. 44 AGAACCGAAATGGAGGAAAGCAAGGC  
SEQ. ID. NO. 42 AGAACTCCTCGGTTATTCTGTTCCTT

SEQ. ID. NO. 44 TCTCTTTAGAACAAATTATCACATAC  
SEQ. ID. NO. 42 AAACAAGAAAGATCTTCTAGAGGAG

SEQ. ID. NO. 44 CCTGGTTTCCAGAACTCCTCGGTTA  
SEQ. ID. NO. 42 AAAATCATGTATTCCCATCTAGTCG

SEQ. ID. NO. 44 TTCTGTTCCTTAAACAAGAAAGATCT  
SEQ. ID. NO. 42 ACTACTTCCCAAGAAATATGATGGACC

SEQ. ID. NO. 44 TCTAGAGGAGAAATCATGTATTCC  
SEQ. ID. NO. 42 CCAGAGAGATGCCCAAGGCAGCCCGA

SEQ. ID. NO. 44 CATCTAGTCGACTACTTCCCAAGAAAT  
SEQ. ID. NO. 42 GAATTCAATTCTGAAGATGTTCTGTGG

SEQ. ID. NO. 44 ATGATGGACCCCAAGAGAGATGCCCA  
SEQ. ID. NO. 42 ACCTGAACCCAGACAGTGACAAAAT

SEQ. ID. NO. 44 GGCAGCCCGAGAAATTCATTCTGAAG  
SEQ. ID. NO. 42 TAACTACTCCCACTTCACTGTGCGCC

SEQ. ID. NO. 44 ATGTTCTGTGGACCTGAACCCAGACA  
SEQ. ID. NO. 42 ACAGACACCGAGAAATATCCGCTTTG

SEQ. ID. NO. 44 GTGACAAAAATTATCTACTCCCACTT  
SEQ. ID. NO. 42 TCTTTGCTGCCGTCAAGGACACCAAT

Figure 13L

SEQ. ID. NO. 44 C A C G T G C G C C A C A G A C A C C G A G A A T  
SEQ. ID. NO. 42 C C T C C A G T T G A A C C T G A A G G G C T G C

SEQ. ID. NO. 44 A T C C G C T T T G T C T T T G C T G C C G T C A  
SEQ. ID. NO. 42 G G T C T G T A C

SEQ. ID. NO. 44 A G G A C A C C A T C C T C C A G T T G A A C C T  
SEQ. ID. NO. 42

SEQ. ID. NO. 44 G A A G G G C T G C G G T C T G T A C  
SEQ. ID. NO. 42

Figure 13M

### ClustalW Formatted Alignments

SEQ. ID. NO. 45 M L L L L L L A P L F L R P P G A G G A Q T P N A  
SEQ. ID. NO. 43 M A S P R S S G Q P G P P P P P P P P A R L L L

SEQ. ID. NO. 45 T S E G C Q I I H P P W E G G I R Y R G L T R D Q  
SEQ. ID. NO. 43 L L L L P L L L P L A P G A W G W A R G A P R P P

SEQ. ID. NO. 45 V K A I N F L P V D Y E I E Y V C R G E R E V V G  
SEQ. ID. NO. 43 P S S P P L S I M G L M P L T K E V A K G S I G R

SEQ. ID. NO. 45 P K V R K C L A N G S W T D M D T P S R C V R I C  
SEQ. ID. NO. 43 G V L P A V E L A I E Q I R N E S L L R P Y F L D

SEQ. ID. NO. 45 S K S Y L T L E N G K V F L T G G D L P A L D G A  
SEQ. ID. NO. 43 L R L Y D T E C D N A K G L K A F Y D A I K Y G P

SEQ. ID. NO. 45 R V D F R C D P D F H L V G S S R S I C S Q G Q W  
SEQ. ID. NO. 43 N H L M V F G G V C P S V T S I I A E S L Q G W N

SEQ. ID. NO. 45 S T P K P H C Q V N R T P H S E R R A V Y I G A L  
SEQ. ID. NO. 43 L V Q L S F A A T T P V L A D K K K Y P Y F F R T

SEQ. ID. NO. 45 F P M S G G W P G G Q A C Q P A V E M A L E D V N  
SEQ. ID. NO. 43 V P S D N A V N P A I L K L L K H Y Q W K R V G T

SEQ. ID. NO. 45 S R R D I L P D Y E L K L I H H D S K C D P G Q A  
SEQ. ID. NO. 43 L T Q D V Q R F S E V R N D L T G V L Y G E D I E

SEQ. ID. NO. 45 T K Y L Y E L L Y N D P I K I I L M P G C S S V S  
SEQ. ID. NO. 43 I S D T E S F S N D P C T S V K K L K G N D V R I

SEQ. ID. NO. 45 T L V A E A A R M W N L I V L S Y G S S S P A L S  
SEQ. ID. NO. 43 I L G Q F D Q N M A A K V F C C A Y E E N M Y G S

SEQ. ID. NO. 45 N R Q R F P T F F R T H P S A T L H N P T R V K L  
SEQ. ID. NO. 43 K Y Q W I I P G W Y E P S W W E Q V H T E A N S S

Figure 14a

SEQ. ID. NO. 45 FEK WGWKKIATIQQTTTEVFTSTLDD  
SEQ. ID. NO. 43 RCLRKNNLLAAMEGYIGVDFEPLSSK

SEQ. ID. NO. 45 LEERVKEAGIEITFRQSFFSDPAVP  
SEQ. ID. NO. 43 QIKTISGKTPQQYEREYNNKRSGVG

SEQ. ID. NO. 45 VKNLKRQDARIIVGLFYETEARKV F  
SEQ. ID. NO. 43 PSKFHGYAYDGIWVIAKTLQRAMET

SEQ. ID. NO. 45 CEVYKERLFGKKYVWFLIGWYADNW  
SEQ. ID. NO. 43 LHASSRHQRIQDFNYTDHTLGRIL

SEQ. ID. NO. 45 FKIYDPSINCTVDEMTEAVEGHITT  
SEQ. ID. NO. 43 NAMNETNPFVGVTGQVVFRNGERMGT

SEQ. ID. NO. 45 EIVMLNPANTRISISNMTSQEFVEKL  
SEQ. ID. NO. 43 IKFTQFQDSREVKVGEYNAVADTLE

SEQ. ID. NO. 45 TKRLKRHPREETGG - FQEAPLAYDAI  
SEQ. ID. NO. 43 IINDTIRFQGSEPPKDKTIILEQLR

SEQ. ID. NO. 45 WALALALNKTSGGGGRSGVRLED FN  
SEQ. ID. NO. 43 KISLPLYSILSALTILGMIMASAF L

SEQ. ID. NO. 45 YNNQTITDQIYRAMNSSSSFEGVSGH  
SEQ. ID. NO. 43 FFNIKNRNQKLIKMS SPYMNLIIL

SEQ. ID. NO. 45 VVFDASGSRMAWTLIEQLQGGSYKK  
SEQ. ID. NO. 43 GGMLSYASIFLFGLDGSFVSEKTFE

SEQ. ID. NO. 45 IGY YDSTKDDL SWSKTDKWIGGSPP  
SEQ. ID. NO. 43 TLCTVRTWILT VGYTTAFGAMFAKT

SEQ. ID. NO. 45 ADQTLVIKTRFLSQKLFISVSVLS  
SEQ. ID. NO. 43 WRVHAIFKNVKMKKKI IKDQKLLVI

SEQ. ID. NO. 45 SLGIVLAVVCLSFNIYN SHVRYIQN  
SEQ. ID. NO. 43 VGGMLLIDLCILICWQAVDPLRRTV

Figure 14b

SEQ. ID. NO. 45 S Q P N L N N L T A V G C S L A L A A V F P L G L  
SEQ. ID. NO. 43 E K Y S M E P D P A G R D I S I R P L L E H C E N

SEQ. ID. NO. 45 D G Y H I G R N Q F P F V C Q A R L W L L G L G F  
SEQ. ID. NO. 43 T H M T I W L G I V Y A Y K G L L M L F G C F L A

SEQ. ID. NO. 45 S L G Y G S M F T K I W W V H T V F T K K E E K K  
SEQ. ID. NO. 43 W E T R N V S I P A L N D S K Y I G M S V Y N V G

SEQ. ID. NO. 45 E W R K T L E P W K L Y A T V G L L V G M D V L T  
SEQ. ID. NO. 43 I M C I I G A A V S F L T R D Q P N V Q F C I V A

SEQ. ID. NO. 45 L A I W Q I V D P L H R T I E T F A K E E P K E D  
SEQ. ID. NO. 43 L V I I F C S T I T L C L V F V P K L I T L R T N

SEQ. ID. NO. 45 I D V S I L P Q L E H C S S R K M N T W L G I F Y  
SEQ. ID. NO. 43 P D A A T Q N R R F Q F T Q N Q K K E D S K T S T

SEQ. ID. NO. 45 G Y K G L L L L L G I F L A Y E T K S V S T E K I  
SEQ. ID. NO. 43 S V T S V N Q A S T S R L E G L Q S E N H R L R M

SEQ. ID. NO. 45 N D H R A V G M A I Y N V A V L C L I T A P V T M  
SEQ. ID. NO. 43 K I T E L D K D L E E V T M Q L Q D T P E K T T Y

SEQ. ID. NO. 45 I L S S Q Q D A A F A F A S L A I V F S S Y I T L  
SEQ. ID. NO. 43 I K Q N H Y Q E L N D I L N L G N F T E S T D G G

SEQ. ID. NO. 45 V V L F V P K M R R L I T R G E W Q S E A Q D T M  
SEQ. ID. NO. 43 K A I L K N H L D Q N P Q L Q W N T T E P S R T C

SEQ. ID. NO. 45 K T G S S T N N N E E E K S R L L E K E N R E L E  
SEQ. ID. NO. 43 K D P I E D I N S P E H I Q R R L S L Q L P I L H

SEQ. ID. NO. 45 K I I A E K E E R V S E L R H Q L Q S R Q Q L R S  
SEQ. ID. NO. 43 H A Y L P S I G G V D A S C V S P C V S P T A S P

SEQ. ID. NO. 45 R R H P P T P P E P S G G L P R G P P E P P D R L  
SEQ. ID. NO. 43 R H R H V P P S F R V M V S G L A A A M T L E S I

Figure 14c

SEQ. ID. NO. 45 SCDGSRVHLLYKAAAMTLESIMACC  
SEQ. ID. NO. 43 MACCLSEEAKEARRINDEIERQLRR

SEQ. ID. NO. 45 LSEEAKEARRINDEIERQLRRDKRD  
SEQ. ID. NO. 43 DKRDARRELKLLLLLGTGESGKSTFI

SEQ. ID. NO. 45 ARRELKLLLLLGTGESGKSTFIKQMR  
SEQ. ID. NO. 43 KQMRIIHGSGYSDEEDKRGFTKLVYQ

SEQ. ID. NO. 45 IIHGSGYSDEEDKRGFTKLVYQNIFT  
SEQ. ID. NO. 43 NIFTAMQAMIRAMD TLKIPYKYEHN

SEQ. ID. NO. 45 AMQAMIRAMD TLKIPYKYEHNKAHA  
SEQ. ID. NO. 43 KAHAQLVREVDVEKVS AFENPYVDA

SEQ. ID. NO. 45 QLVREVDVEKVS AFENPYVDAIKSL  
SEQ. ID. NO. 43 IKS LWN DPGIQECYDRRREYQLSDS

SEQ. ID. NO. 45 WNDPGIQECYDRRREYQLSDSTKYY  
SEQ. ID. NO. 43 TKYYLNDLDRVADPAYLPTQQQDVLR

SEQ. ID. NO. 45 LNDLDRVADPAYLPTQQQDVLRVRVP  
SEQ. ID. NO. 43 VRVP TTGII EYPFDLQSVIFRMVDV

SEQ. ID. NO. 45 TTGII EYPFDLQSVIFRMVDVGGQR  
SEQ. ID. NO. 43 GGQRSE R R KWIHC FENVTSIMFLVA

SEQ. ID. NO. 45 SE R R KWIHC FENVTSIMFLVALSEY  
SEQ. ID. NO. 43 LSEYDQVLVESDNENRMEESKALFR

SEQ. ID. NO. 45 DQVLVESDNENRMEESKALFR TIIT  
SEQ. ID. NO. 43 TIITYPWFQNSSSVILFLNKKDLLEE

SEQ. ID. NO. 45 YPWFQNSSSVILFLNKKDLLEEKIMY  
SEQ. ID. NO. 43 KIMYSHLVDFPEYDGPQRDAQAAR

SEQ. ID. NO. 45 SHLVDFPEYDGPQRDAQAAREFIL  
SEQ. ID. NO. 43 EFILKMFVDLNPDS DKINYSHF TCA

Figure 14d

SEQ. ID. NO. 45 K M F V D L N P D S D K I I Y S H F T C A T D T E  
SEQ. ID. NO. 43 T D T E N I R F V F A A V K D T I L Q L N L K G C

SEQ. ID. NO. 45 N I R F V F A A V K D T I L Q L N L K G C G L Y  
SEQ. ID. NO. 43 G L Y

Figure 14e

mGluR8//CaR\*Gqi5

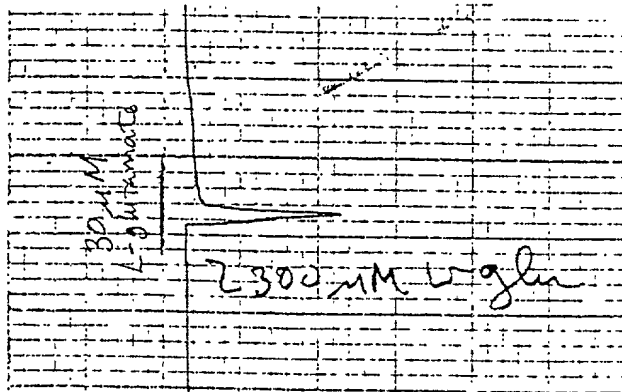
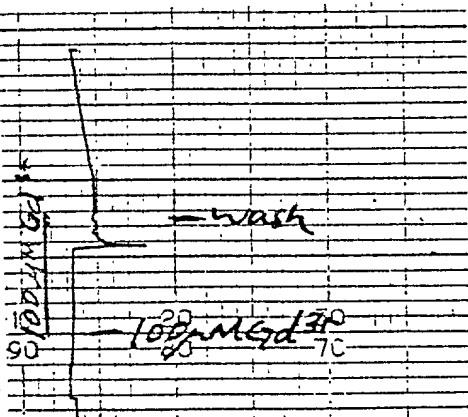


Figure 15

CaR/mGluR2\*Gqi5



mGluR2//CaR\*Gqi5

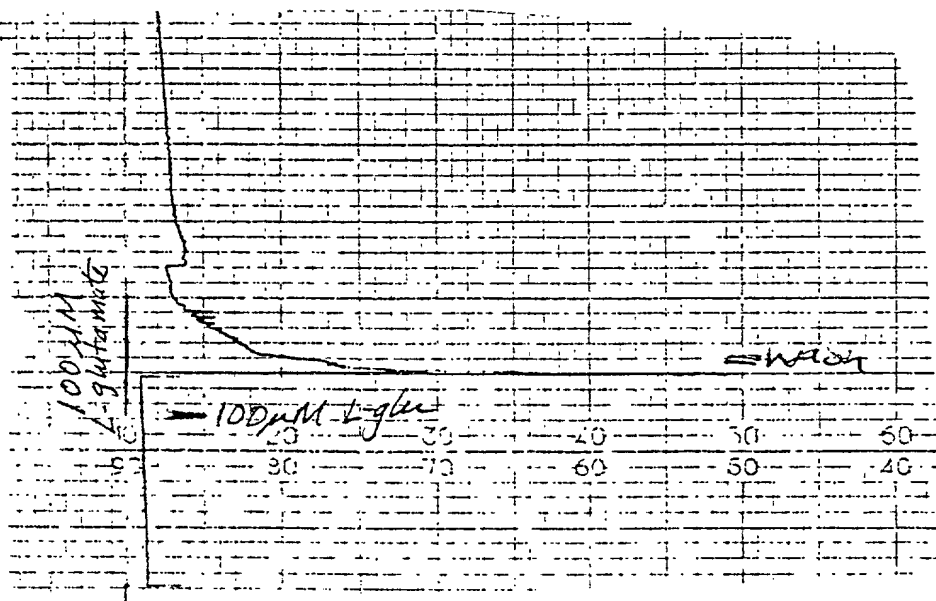




FIGURE 16A

SEQ. ID. NO. 48 M V C E G K R S A S C P C F F L L T A K F Y W I L T M M Q R  
 SEQ. ID. NO. 49 M V C E G K R S A S C P C F F L L T A K F Y W I L T M M Q R  
 SEQ. ID. NO. 50 M V C E G K R S A S C P C F F L L T A K F Y W I L T M M Q R

SEQ. ID. NO. 48 T H S Q E Y A H S I R I D G D I T L G G L F P V H G R G S E  
 SEQ. ID. NO. 49 T H S Q E Y A H S I R I D G D I T L G G L F P V H G R G S E  
 SEQ. ID. NO. 50 T H S Q E Y A H S I R V D G D I I L G G L F P V H A K G E R

SEQ. ID. NO. 48 G K P C G E L K K E K G I H R L E A M L F A L D R I N N D P  
 SEQ. ID. NO. 49 G K P C G E L K K E K G I H R L E A M L F A L D R I N N D P  
 SEQ. ID. NO. 50 G V P C G E L K K E K G I H R L E A M L Y A I D Q I N K D P

SEQ. ID. NO. 48 D L L P N I T L G A R I L D T C S R D T H A L E Q S L T F V  
 SEQ. ID. NO. 49 D L L P N I T L G A R I L D T C S R D T H A L E Q S L T F V  
 SEQ. ID. NO. 50 D L L S N I T L G V R I L D T C S R D T Y A L E Q S L T F V

SEQ. ID. NO. 48 Q A L I E K D G T E V R C G S G G P P I I T K P E R V V G V  
 SEQ. ID. NO. 49 Q A L I E K D G T E V R C G S G G P P I I T K P E R V V G V  
 SEQ. ID. NO. 50 Q A L I E K D A S D V K C A N G D P P I F T K P D K I S G V

SEQ. ID. NO. 48 I G A S G S S V S I M V A N I L R L F K I P Q I S Y A S T A  
 SEQ. ID. NO. 49 I G A S G S S V S I M V A N I L R L F K I P Q I S Y A S T A  
 SEQ. ID. NO. 50 I G A A A S S V S I M V A N I L R L F K I P Q I S Y A S T A

SEQ. ID. NO. 48 P D L S D N S R Y D F F S R V V P S D T Y Q A Q A M V D I V  
 SEQ. ID. NO. 49 P D L S D N S R Y D F F S R V V P S D T Y Q A Q A M V D I V  
 SEQ. ID. NO. 50 P E L S D N T R Y D F F S R V V P P D S Y Q A Q A M V D I V

SEQ. ID. NO. 48 R A L K W N Y V S T V A S E G S Y G E S G V E A F I Q K S R  
 SEQ. ID. NO. 49 R A L K W N Y V S T V A S E G S Y G E S G V E A F I Q K S R  
 SEQ. ID. NO. 50 T A L G W N Y V S T L A S E G N Y G E S G V E A F T Q I S R

SEQ. ID. NO. 48 E D G G V C I A Q S V K I P R E P K A G E F D K I I R R L L  
 SEQ. ID. NO. 49 E D G G V C I A Q S V K I P R E P K A G E F D K I I R R L L  
 SEQ. ID. NO. 50 E I G G V C I A Q S Q K I P R E P R P G E F E K I I K R L L

FIGURE 16B

SEQ. ID. NO. 48 E T S N A R A V I I F A N E D D I R R V L E A A R R A N Q T  
 SEQ. ID. NO. 49 E T S N A R A V I I F A N E D D I R R V L E A A R R A N Q T  
 SEQ. ID. NO. 50 E T P N A R A V I M F A N E D D I R R I L E A A K K L N Q S

SEQ. ID. NO. 48 G H F F W M G S D S W G S K I A P V L H L E E V A E G A V T  
 SEQ. ID. NO. 49 G H F F W M G S D S W G S K I A P V L H L E E V A E G A V T  
 SEQ. ID. NO. 50 G H F L W I G S D S W G S K I A P V Y Q Q E E I A E G A V T

SEQ. ID. NO. 48 I L P K R M S V R G F D R Y F S S R T L D N N R R N I W F A  
 SEQ. ID. NO. 49 I L P K R M S V R G F D R Y F S S R T L D N N R R N I W F A  
 SEQ. ID. NO. 50 I L P K R A S I D G F D R Y F R S R T L A N N R R N V W F A

SEQ. ID. NO. 48 E F W E D N F H C K L S R H A L K K G S H V K K C T N R E R  
 SEQ. ID. NO. 49 E F W E D N F H C K L S R H A L K K G S H V K K C T N R E R  
 SEQ. ID. NO. 50 E F W E E N F G C K L G S H G K R N - S H I K K C T G L E R

SEQ. ID. NO. 48 I G Q D S A Y E Q E G K V Q F V I D A V Y A M G H A L H A M  
 SEQ. ID. NO. 49 I G Q D S A Y E Q E G K V Q F V I D A V Y A M G H A L H A M  
 SEQ. ID. NO. 50 I A R D S S Y E Q E G K V Q F V I D A V Y S M A Y A L H N M

SEQ. ID. NO. 48 H R D L C P G R V G L C P R M D P V D G T Q L L K Y I R N V  
 SEQ. ID. NO. 49 H R D L C P G R V G L C P R M D P V D G T Q L L K Y I R N V  
 SEQ. ID. NO. 50 H K D L C P G Y I G L C P R M S T I D G K E L L G Y I R A V

SEQ. ID. NO. 48 N F S G I A G N P V T F N E N G D A P G R Y D I Y Q Y Q L R  
 SEQ. ID. NO. 49 N F S G I A G N P V T F N E N G D A P G R Y D I Y Q Y Q L R  
 SEQ. ID. NO. 50 N F N G S A G T P V T F N E N G D A P G R Y D I F Q Y Q I T

SEQ. ID. NO. 48 N D S A E Y K V I G S W T D H L H L R I E R M H W P G S G Q  
 SEQ. ID. NO. 49 N D S A E Y K V I G S W T D H L H L R I E R M H W P G S G Q  
 SEQ. ID. NO. 50 N K S T E Y K V I G H W T N Q L H L K V E D M Q W A H R E H

SEQ. ID. NO. 48 Q L P R S I C S L P C Q P G E R K K T V K G M P C C W H C E  
 SEQ. ID. NO. 49 Q L P R S I C S L P C Q P G E R K K T V K G M P C C W H C E  
 SEQ. ID. NO. 50 T H P A S V C S L P C K P G E R K K T V K G V P C C W H C E

SEQ. ID. NO. 48 P C T G Y Q Y Q V D R Y T C K T C P Y D M R P T E N R T G C  
 SEQ. ID. NO. 49 P C T G Y Q Y Q V D R Y T C K T C P Y D M R P T E N R T G C  
 SEQ. ID. NO. 50 R C E G Y N Y Q V D E L S C E L C P L D Q R P N M N R T G C

FIGURE 16C

SEQ. ID. NO. 48 R P I P I I K L E W G S P W A V L P L F L A V V G I A A T L  
 SEQ. ID. NO. 49 R P I P I I K L E W G S P W A V L P L F L A V V G I A A T L  
 SEQ. ID. NO. 50 Q L I P I I K L E W H S P W A V V P V F V A I L G I I A T T

SEQ. ID. NO. 48 F V V I T F V R Y N D T P I V K A S G R E L S Y V L L A G I  
 SEQ. ID. NO. 49 F V V I T F V R Y N D T P I V K A S G R E L S Y V L L A G I  
 SEQ. ID. NO. 50 F V I V T F V R Y N D T P I V R A S G R E L S Y V L L T G I

SEQ. ID. NO. 48 F L C Y A T T F L M I A E P D L G T C S L R R I F L G L G M  
 SEQ. ID. NO. 49 F L C Y A T T F L M I A E P D L G T C S L R R I F L G L G M  
 SEQ. ID. NO. 50 F L C Y S I T F L M I A A P D T I I C S F R R V F L G L G M

SEQ. ID. NO. 48 S I S Y A A L L T K T N R I Y R I F E Q G K R S V S A P R F  
 SEQ. ID. NO. 49 S I S Y A A L L T K T N R I Y R I F E Q G K R S V S A P R F  
 SEQ. ID. NO. 50 C F S Y A A L L T K T N R I H R I F E Q G K K S V T A P K F

SEQ. ID. NO. 48 I S P A S Q L A I T F S L I S L Q L L G I C V W F V V D P S  
 SEQ. ID. NO. 49 I S P A S Q L A I T F S L I S L Q L L G I C V W F V V D P S  
 SEQ. ID. NO. 50 I S P A S Q L V I T F S L I S V Q L L G V F V W F V V D P P

SEQ. ID. NO. 48 H S V V D F Q D Q R T L D P R F A R G V L K C D I S D L S L  
 SEQ. ID. NO. 49 H S V V D F Q D Q R T L D P R F A R G V L K C D I S D L S L  
 SEQ. ID. NO. 50 H I I I D Y G E Q R T L D P E K A R G V L K C D I S D L S L

SEQ. ID. NO. 48 I C L L G Y S M L L M V T C T V Y A I K T R G V P E T F N E  
 SEQ. ID. NO. 49 I C L L G Y S M L L M V T C T V Y A I K T R G V P E T F N E  
 SEQ. ID. NO. 50 I C S L G Y S I L L M V T C T V Y A I K T R G V P E T F N E

SEQ. ID. NO. 48 A K P I G F T M Y T T C I V W L A F I P I F F G T S Q S A D  
 SEQ. ID. NO. 49 A K P I G F T M Y T T C I V W L A F I P I F F G T S Q S A D  
 SEQ. ID. NO. 50 A K P I G F T M Y T T C I I W L A F I P I F F G T A Q S A E

SEQ. ID. NO. 48 K L Y I Q T T T L T V S V S L S A S V S L G M L Y M P K V Y  
 SEQ. ID. NO. 49 K L Y I Q T T T L T V S V S L S A S V S L G M L Y M P K V Y  
 SEQ. ID. NO. 50 K M Y I Q T T T L T V S M S L S A S V S L G M L Y M P K V Y

SEQ. ID. NO. 48 I I L F H P E Q N V P K R K R S L K A V V T A A T M S N K F  
 SEQ. ID. NO. 49 I I L F H P E Q N T I E E V R C S T A A H A F K V A A R A T  
 SEQ. ID. NO. 50 I I I F H P E Q N T I E E V R C S T A A H A F K V A A R A T

FIGURE 16D

SEQ. ID. NO. 48 T Q K G N F R P N G E A K S E L C E N L E A P A L A T K Q T  
SEQ. ID. NO. 49 L R R S N V S R K R S S S L G G S T G S T P S S S I S S K S  
SEQ. ID. NO. 50 L R R S N V S R K R S S S L G G S T G S T P S S S I S S K S

SEQ. ID. NO. 48 Y V T Y T N H A I  
SEQ. ID. NO. 49 N S E D P F P Q P E R Q K Q Q Q P L A L T Q Q E Q Q Q Q P L  
SEQ. ID. NO. 50 N S E D P F P Q P E R Q K Q Q Q P L A L T Q Q E Q Q Q Q P L

SEQ. ID. NO. 48  
SEQ. ID. NO. 49 T L P Q Q Q R S Q Q Q P R C K Q K V I F G S G T V T F S L S  
SEQ. ID. NO. 50 T L P Q Q Q R S Q Q Q P R C K Q K V I F G S G T V T F S L S

SEQ. ID. NO. 48  
SEQ. ID. NO. 49 F D E P Q K N A M A H G N S T H Q N S L E A Q K S S D T L T  
SEQ. ID. NO. 50 F D E P Q K N A M A H G N S T H Q N S L E A Q K S S D T L T

SEQ. ID. NO. 48  
SEQ. ID. NO. 49 R H Q P L L P L Q C G E T D L D L T V Q E T G L Q G P V G G  
SEQ. ID. NO. 50 R H Q P L L P L Q C G E T D L D L T V Q E T G L Q G P V G G

SEQ. ID. NO. 48  
SEQ. ID. NO. 49 D Q R P E V E D P E E L S P A L V V S S S Q S F V I S G G G  
SEQ. ID. NO. 50 D Q R P E V E D P E E L S P A L V V S S S Q S F V I S G G G

SEQ. ID. NO. 48  
SEQ. ID. NO. 49 S T V T E N V V N S A A A M T L E S I M A C C L S E E A K E  
SEQ. ID. NO. 50 S T V T E N V V N S A A A M T L E S I M A C C L S E E A K E

SEQ. ID. NO. 48  
SEQ. ID. NO. 49 A R R I N D E I E R Q L R R D K R D A R R E L K L L L L G T  
SEQ. ID. NO. 50 A R R I N D E I E R Q L R R D K R D A R R E L K L L L L G T

SEQ. ID. NO. 48  
SEQ. ID. NO. 49 G E S G K S T F I K Q M R I I H G S G Y S D E D K R G F T K  
SEQ. ID. NO. 50 G E S G K S T F I K Q M R I I H G S G Y S D E D K R G F T K

SEQ. ID. NO. 48  
SEQ. ID. NO. 49 L V Y Q N I F T A M Q A M I R A M D T L K I P Y K Y E H N K  
SEQ. ID. NO. 50 L V Y Q N I F T A M Q A M I R A M D T L K I P Y K Y E H N K

FIGURE 16E

SEQ. ID. NO. 48

SEQ. ID. NO. 49

SEQ. ID. NO. 50

A H A Q L V R E V D V E K V S A F E N P Y V D A I K S L W N  
A H A Q L V R E V D V E K V S A F E N P Y V D A I K S L W N

SEQ. ID. NO. 48

SEQ. ID. NO. 49

SEQ. ID. NO. 50

D P G I Q E C Y D R R R E Y Q L S D S T K Y Y L N D L D R V  
D P G I Q E C Y D R R R E Y Q L S D S T K Y Y L N D L D R V

SEQ. ID. NO. 48

SEQ. ID. NO. 49

SEQ. ID. NO. 50

A D P A Y L P T Q Q D V L R V R V P T T G I I E Y P F D L Q  
A D P A Y L P T Q Q D V L R V R V P T T G I I E Y P F D L Q

SEQ. ID. NO. 48

SEQ. ID. NO. 49

SEQ. ID. NO. 50

S V I F R M V D V G G Q R S E R R K W I H C F E N V T S I M  
S V I F R M V D V G G Q R S E R R K W I H C F E N V T S I M

SEQ. ID. NO. 48

SEQ. ID. NO. 49

SEQ. ID. NO. 50

F L V A L S E Y D Q V L V E S D N E N R M E E S K A L F R T  
F L V A L S E Y D Q V L V E S D N E N R M E E S K A L F R T

SEQ. ID. NO. 48

SEQ. ID. NO. 49

SEQ. ID. NO. 50

I I T Y P W F Q N S S V I L F L N K K D L L E E K I M Y S H  
I I T Y P W F Q N S S V I L F L N K K D L L E E K I M Y S H

SEQ. ID. NO. 48

SEQ. ID. NO. 49

SEQ. ID. NO. 50

L V D Y F P E Y D G P Q R D A Q A A R E F I L K M F V D L N  
L V D Y F P E Y D G P Q R D A Q A A R E F I L K M F V D L N

SEQ. ID. NO. 48

SEQ. ID. NO. 49

SEQ. ID. NO. 50

P D S D K I I Y S H F T C A T D T E N I R F V F A A V K D T  
P D S D K I I Y S H F T C A T D T E N I R F V F A A V K D T

SEQ. ID. NO. 48

SEQ. ID. NO. 49

SEQ. ID. NO. 50

I L Q L N L K D C G L F  
I L Q L N L K D C G L F

## SEQUENCE LISTING

<110> NPS PHARMACEUTICALS, INC.

<120> G-PROTEIN FUSION RECEPTORS AND CHIMERIC  
GABAB RECEPTORS

<130> 241/086-CIP

<140> TO BE ASSIGNED

<141> HEREWITH

<150> US 60/080,671

<151> 1998-04-03

<150> PCT/US99/07333

<151> 1999-04-02

<160> 50

<170> FastSEQ for Windows Version 3.0

<210> 1

<211> 612

<212> PRT

<213> Human

<400> 1

```

Met Ala Phe Tyr Ser Cys Cys Trp Val Leu Leu Ala Leu Thr Trp His
 1              5              10              15

Thr Ser Ala Tyr Gly Pro Asp Gln Arg Ala Gln Lys Lys Gly Asp Ile
      20              25              30

Ile Leu Gly Gly Leu Phe Pro Ile His Phe Gly Val Ala Ala Lys Asp
 35              40              45

Gln Asp Leu Lys Ser Arg Pro Glu Ser Val Glu Cys Ile Arg Tyr Asn
 50              55              60

Phe Arg Gly Phe Arg Trp Leu Gln Ala Met Ile Phe Ala Ile Glu Glu
 65              70              75              80

Ile Asn Ser Ser Pro Ala Leu Leu Pro Asn Leu Thr Leu Gly Tyr Arg
      85              90              95

Ile Phe Asp Thr Cys Asn Thr Val Ser Lys Ala Leu Glu Ala Thr Leu
 100              105              110

```

Ser Phe Val Ala Gln Asn Lys Ile Asp Ser Leu Asn Leu Asp Glu Phe  
 115 120 125

Cys Asn Cys Ser Glu His Ile Pro Ser Thr Ile Ala Val Val Gly Ala  
 130 135 140

Thr Gly Ser Gly Val Ser Thr Ala Val Ala Asn Leu Leu Gly Leu Phe  
 145 150 155 160

Tyr Ile Pro Gln Val Ser Tyr Ala Ser Ser Ser Arg Leu Leu Ser Asn  
 165 170 175

Lys Asn Gln Phe Lys Ser Phe Leu Arg Thr Ile Pro Asn Asp Glu His  
 180 185 190

Gln Ala Thr Ala Met Ala Asp Ile Ile Glu Tyr Phe Arg Trp Asn Trp  
 195 200 205

Val Gly Thr Ile Ala Ala Asp Asp Asp Tyr Gly Arg Pro Gly Ile Glu  
 210 215 220

Lys Phe Arg Glu Glu Ala Glu Glu Arg Asp Ile Cys Ile Asp Phe Ser  
 225 230 235 240

Glu Leu Ile Ser Gln Tyr Ser Asp Glu Glu Glu Ile Gln His Val Val  
 245 250 255

Glu Val Ile Gln Asn Ser Thr Ala Lys Val Ile Val Val Phe Ser Ser  
 260 265 270

Gly Pro Asp Leu Glu Pro Leu Ile Lys Glu Ile Val Arg Arg Asn Ile  
 275 280 285

Thr Gly Lys Ile Trp Leu Ala Ser Glu Ala Trp Ala Ser Ser Ser Leu  
 290 295 300

Ile Ala Met Pro Gln Tyr Phe His Val Val Gly Gly Thr Ile Gly Phe  
 305 310 315 320

Ala Leu Lys Ala Gly Gln Ile Pro Gly Phe Arg Glu Phe Leu Lys Lys  
 325 330 335

Val His Pro Arg Lys Ser Val His Asn Gly Phe Ala Lys Glu Phe Trp  
 340 345 350

Glu Glu Thr Phe Asn Cys His Leu Gln Glu Gly Ala Lys Gly Pro Leu  
 355 360 365

Pro Val Asp Thr Phe Leu Arg Gly His Glu Glu Ser Gly Asp Arg Phe  
 370 375 380

Ser Asn Ser Ser Thr Ala Phe Arg Pro Leu Cys Thr Gly Asp Glu Asn  
 385 390 395 400

Ile Ser Ser Val Glu Thr Pro Tyr Ile Asp Tyr Thr His Leu Arg Ile

405 410 415  
 Ser Tyr Asn Val Tyr Leu Ala Val Tyr Ser Ile Ala His Ala Leu Gln  
 420 425 430  
 Asp Ile Tyr Thr Cys Leu Pro Gly Arg Gly Leu Phe Thr Asn Gly Ser  
 435 440 445  
 Cys Ala Asp Ile Lys Lys Val Glu Ala Trp Gln Val Leu Lys His Leu  
 450 455 460  
 Arg His Leu Asn Phe Thr Asn Asn Met Gly Glu Gln Val Thr Phe Asp  
 465 470 475 480  
 Glu Cys Gly Asp Leu Val Gly Asn Tyr Ser Ile Ile Asn Trp His Leu  
 485 490 495  
 Ser Pro Glu Asp Gly Ser Ile Val Phe Lys Glu Val Gly Tyr Tyr Asn  
 500 505 510  
 Val Tyr Ala Lys Lys Gly Glu Arg Leu Phe Ile Asn Glu Glu Lys Ile  
 515 520 525  
 Leu Trp Ser Gly Phe Ser Arg Glu Val Pro Phe Ser Asn Cys Ser Arg  
 530 535 540  
 Asp Cys Leu Ala Gly Thr Arg Lys Gly Ile Ile Glu Gly Glu Pro Thr  
 545 550 555 560  
 Cys Cys Phe Glu Cys Val Glu Cys Pro Asp Gly Glu Tyr Ser Asp Glu  
 565 570 575  
 Thr Asp Ala Ser Ala Cys Asn Lys Cys Pro Asp Asp Phe Trp Ser Asn  
 580 585 590  
 Glu Asn His Thr Ser Cys Ile Ala Lys Glu Ile Glu Phe Leu Ser Trp  
 595 600 605  
 Thr Glu Pro Phe  
 610

<210> 2  
 <211> 590  
 <212> PRT  
 <213> Human

<400> 2

Met Leu Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly  
 1 5 10 15  
 Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile  
 20 25 30



Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg  
 35 40 45  
 Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu  
 50 55 60  
 Tyr Val Cys Arg Gly Glu Arg Glu Val Val Gly Pro Lys Val Arg Lys  
 65 70 75 80  
 Cys Leu Ala Asn Gly Ser Trp Thr Asp Met Asp Thr Pro Ser Arg Cys  
 85 90 95  
 Val Arg Ile Cys Ser Lys Ser Tyr Leu Thr Leu Glu Asn Gly Lys Val  
 100 105 110  
 Phe Leu Thr Gly Gly Asp Leu Pro Ala Leu Asp Gly Ala Arg Val Asp  
 115 120 125  
 Phe Arg Cys Asp Pro Asp Phe His Leu Val Gly Ser Ser Arg Ser Ile  
 130 135 140  
 Cys Ser Gln Gly Gln Trp Ser Thr Pro Lys Pro His Cys Gln Val Asn  
 145 150 155 160  
 Arg Thr Pro His Ser Glu Arg Arg Ala Val Tyr Ile Gly Ala Leu Phe  
 165 170 175  
 Pro Met Ser Gly Gly Trp Pro Gly Gly Gln Ala Cys Gln Pro Ala Val  
 180 185 190  
 Glu Met Ala Leu Glu Asp Val Asn Ser Arg Arg Asp Ile Leu Pro Asp  
 195 200 205  
 Tyr Glu Leu Lys Leu Ile His His Asp Ser Lys Cys Asp Pro Gly Gln  
 210 215 220  
 Ala Thr Lys Tyr Leu Tyr Glu Leu Leu Tyr Asn Asp Pro Ile Lys Ile  
 225 230 235 240  
 Ile Leu Met Pro Gly Cys Ser Ser Val Ser Thr Leu Val Ala Glu Ala  
 245 250 255  
 Ala Arg Met Trp Asn Leu Ile Val Leu Ser Tyr Gly Ser Ser Ser Pro  
 260 265 270  
 Ala Leu Ser Asn Arg Gln Arg Phe Pro Thr Phe Phe Arg Thr His Pro  
 275 280 285  
 Ser Ala Thr Leu His Asn Pro Thr Arg Val Lys Leu Phe Glu Lys Trp  
 290 295 300  
 Gly Trp Lys Lys Ile Ala Thr Ile Gln Gln Thr Thr Glu Val Phe Thr  
 305 310 315 320  
 Ser Thr Leu Asp Asp Leu Glu Glu Arg Val Lys Glu Ala Gly Ile Glu  
 325 330 335

Ile Thr Phe Arg Gln Ser Phe Phe Ser Asp Pro Ala Val Pro Val Lys  
 340 345 350  
 Asn Leu Lys Arg Gln Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu  
 355 360 365  
 Thr Glu Ala Arg Lys Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe  
 370 375 380  
 Gly Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp  
 385 390 395 400  
 Phe Lys Ile Tyr Asp Pro Ser Ile Asn Cys Thr Val Asp Glu Met Thr  
 405 410 415  
 Glu Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro  
 420 425 430  
 Ala Asn Thr Arg Ser Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu  
 435 440 445  
 Lys Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe  
 450 455 460  
 Gln Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala  
 465 470 475 480  
 Leu Asn Lys Thr Ser Gly Gly Gly Gly Arg Ser Gly Val Arg Leu Glu  
 485 490 495  
 Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala  
 500 505 510  
 Met Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp  
 515 520 525  
 Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly  
 530 535 540  
 Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu  
 545 550 555 560  
 Ser Trp Ser Lys Thr Asp Lys Trp Ile Gly Gly Ser Pro Pro Ala Asp  
 565 570 575  
 Gln Thr Leu Val Ile Lys Thr Phe Arg Phe Leu Ser Gln Lys  
 580 585 590

<210> 3  
 <211> 473  
 <212> PRT  
 <213> Human

<400> 3

```

Met Gly Pro Gly Ala Pro Phe Ala Arg Val Gly Trp Pro Leu Pro Leu
 1           5           10           15

Leu Val Val Met Ala Ala Gly Val Ala Pro Val Trp Ala Ser His Ser
      20           25           30

Pro His Leu Pro Arg Pro His Ser Arg Val Pro Pro His Pro Ser Ser
      35           40           45

Glu Arg Arg Ala Val Tyr Ile Gly Ala Leu Phe Pro Met Ser Gly Gly
 50           55           60

Trp Pro Gly Gly Gln Ala Cys Gln Pro Ala Val Glu Met Ala Leu Glu
65           70           75           80

Asp Val Asn Ser Arg Arg Asp Ile Leu Pro Asp Tyr Glu Leu Lys Leu
      85           90           95

Ile His His Asp Ser Lys Cys Asp Pro Gly Gln Ala Thr Lys Tyr Leu
      100          105          110

Tyr Glu Leu Leu Tyr Asn Asp Pro Ile Lys Ile Ile Leu Met Pro Gly
      115          120          125

Cys Ser Ser Val Ser Thr Leu Val Ala Glu Ala Ala Arg Met Trp Asn
      130          135          140

Leu Ile Val Leu Ser Tyr Gly Ser Ser Ser Pro Ala Leu Ser Asn Arg
      145          150          155          160

Gln Arg Phe Pro Thr Phe Phe Arg Thr His Pro Ser Ala Thr Leu His
      165          170          175

Asn Pro Thr Arg Val Lys Leu Phe Glu Lys Trp Gly Trp Lys Lys Ile
      180          185          190

Ala Thr Ile Gln Gln Thr Thr Glu Val Phe Thr Ser Thr Leu Asp Asp
      195          200          205

Leu Glu Glu Arg Val Lys Glu Ala Gly Ile Glu Ile Thr Phe Arg Gln
      210          215          220

Ser Phe Phe Ser Asp Pro Ala Val Pro Val Lys Asn Leu Lys Arg Gln
      225          230          235          240

Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu Thr Glu Ala Arg Lys
      245          250          255

Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe Gly Lys Lys Tyr Val
      260          265          270

```

Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp Phe Lys Ile Tyr Asp  
 275 280 285  
 Pro Ser Ile Asn Cys Thr Val Asp Glu Met Thr Glu Ala Val Glu Gly  
 290 295 300  
 His Ile Thr Thr Glu Ile Val Met Leu Asn Pro Ala Asn Thr Arg Ser  
 305 310 315 320  
 Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu Lys Leu Thr Lys Arg  
 325 330 335  
 Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln Glu Ala Pro Leu  
 340 345 350  
 Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala Leu Asn Lys Thr Ser  
 355 360 365  
 Gly Gly Gly Gly Arg Ser Gly Val Arg Leu Glu Asp Phe Asn Tyr Asn  
 370 375 380  
 Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met Asn Ser Ser Ser  
 385 390 395 400  
 Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala Ser Gly Ser Arg  
 405 410 415  
 Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly Ser Tyr Lys Lys  
 420 425 430  
 Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser Trp Ser Lys Thr  
 435 440 445  
 Asp Lys Trp Ile Gly Gly Ser Pro Pro Ala Asp Gln Thr Leu Val Ile  
 450 455 460  
 Lys Thr Phe Arg Phe Leu Ser Gln Lys  
 465 470

<210> 4  
 <211> 480  
 <212> PRT  
 <213> Human

<400> 4

Met Ala Ser Pro Arg Ser Ser Gly Gln Pro Gly Pro Xaa Pro Pro Pro  
 1 5 10 15  
 Pro Pro Pro Pro Ala Arg Leu Leu Leu Leu Leu Leu Leu Pro Leu Leu  
 20 25 30  
 Leu Pro Leu Ala Pro Gly Ala Trp Gly Trp Ala Arg Gly Ala Pro Arg

35					40					45					
Pro	Pro	Pro	Ser	Ser	Pro	Pro	Leu	Ser	Ile	Met	Gly	Leu	Met	Pro	Leu
50					55					60					
Thr	Lys	Glu	Val	Ala	Lys	Gly	Ser	Ile	Gly	Arg	Gly	Val	Leu	Pro	Ala
65					70					75					80
Val	Glu	Leu	Ala	Ile	Glu	Gln	Ile	Arg	Asn	Glu	Ser	Leu	Leu	Arg	Pro
				85					90					95	
Tyr	Phe	Leu	Asp	Leu	Arg	Leu	Tyr	Asp	Thr	Glu	Cys	Asp	Asn	Ala	Lys
			100					105					110		
Gly	Leu	Lys	Ala	Phe	Tyr	Asp	Ala	Ile	Lys	Tyr	Gly	Pro	Asn	His	Leu
		115					120					125			
Met	Val	Phe	Gly	Gly	Val	Cys	Pro	Ser	Val	Thr	Ser	Ile	Ile	Ala	Glu
	130					135						140			
Ser	Leu	Gln	Gly	Trp	Asn	Leu	Val	Gln	Leu	Ser	Phe	Ala	Ala	Thr	Thr
145					150					155					160
Pro	Val	Leu	Ala	Asp	Lys	Lys	Lys	Tyr	Pro	Tyr	Phe	Phe	Arg	Thr	Val
				165					170					175	
Pro	Ser	Asp	Asn	Ala	Val	Asn	Pro	Ala	Ile	Leu	Lys	Leu	Leu	Lys	His
			180					185					190		
Tyr	Gln	Trp	Lys	Arg	Val	Gly	Thr	Leu	Thr	Gln	Asp	Val	Gln	Arg	Phe
		195					200					205			
Ser	Glu	Val	Arg	Asn	Asp	Leu	Thr	Gly	Val	Leu	Tyr	Gly	Glu	Asp	Ile
	210					215					220				
Glu	Ile	Ser	Asp	Thr	Glu	Ser	Phe	Ser	Asn	Asp	Pro	Cys	Thr	Ser	Val
225					230					235					240
Lys	Lys	Leu	Lys	Gly	Asn	Asp	Val	Arg	Ile	Ile	Leu	Gly	Gln	Phe	Asp
				245					250					255	
Gln	Asn	Met	Ala	Ala	Lys	Val	Phe	Cys	Cys	Ala	Tyr	Glu	Glu	Asn	Met
			260					265					270		
Tyr	Gly	Ser	Lys	Tyr	Gln	Trp	Ile	Ile	Pro	Gly	Trp	Tyr	Glu	Pro	Ser
		275					280					285			
Trp	Trp	Glu	Gln	Val	His	Thr	Glu	Ala	Asn	Ser	Ser	Arg	Cys	Leu	Arg
		290					295					300			
Lys	Asn	Leu	Leu	Ala	Ala	Met	Glu	Gly	Tyr	Ile	Gly	Val	Asp	Phe	Glu
305					310					315					320
Pro	Leu	Ser	Ser	Lys	Gln	Ile	Lys	Thr	Ile	Ser	Gly	Lys	Thr	Pro	Gln
				325					330					335	

Gln Tyr Glu Arg Glu Tyr Asn Asn Lys Arg Ser Gly Val Gly Pro Ser  
340 345 350

Lys Phe His Gly Tyr Ala Tyr Asp Gly Ile Trp Val Ile Ala Lys Thr  
355 360 365

Leu Gln Arg Ala Met Glu Thr Leu His Ala Ser Ser Arg His Gln Arg  
370 375 380

Ile Gln Asp Phe Asn Tyr Thr Asp His Thr Leu Gly Arg Ile Ile Leu  
385 390 395 400

Asn Ala Met Asn Glu Thr Asn Phe Phe Gly Val Thr Gly Gln Val Val  
405 410 415

Phe Arg Asn Gly Glu Arg Met Gly Thr Ile Lys Phe Thr Gln Phe Gln  
420 425 430

Asp Ser Arg Glu Val Lys Val Gly Glu Tyr Asn Ala Val Ala Asp Thr  
435 440 445

Leu Glu Ile Ile Asn Asp Thr Ile Arg Phe Gln Gly Ser Glu Pro Pro  
450 455 460

Lys Asp Lys Thr Ile Ile Leu Glu Gln Leu Arg Lys Ile Ser Leu Pro  
465 470 475 480

<210> 5

<211> 583

<212> PRT

<213> Human

<400> 5

Met Val Cys Glu Gly Lys Arg Ser Ala Ser Cys Pro Cys Phe Phe Leu  
1 5 10 15

Leu Thr Ala Lys Phe Tyr Trp Ile Leu Thr Met Met Gln Arg Thr His  
20 25 30

Ser Gln Glu Tyr Ala His Ser Ile Arg Val Asp Gly Asp Ile Ile Leu  
35 40 45

Gly Gly Leu Phe Pro Val His Ala Lys Gly Glu Arg Gly Val Pro Cys  
50 55 60

Gly Glu Leu Lys Lys Glu Lys Gly Ile His Arg Leu Glu Ala Met Leu  
65 70 75 80

Tyr Ala Ile Asp Gln Ile Asn Lys Asp Pro Asp Leu Leu Ser Asn Ile  
85 90 95

Thr Leu Gly Val Arg Ile Leu Asp Thr Cys Ser Arg Asp Thr Tyr Ala  
100 105 110

11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47 48 49 50 51 52 53 54 55 56 57 58 59 60 61 62 63 64 65 66 67 68 69 70 71 72 73 74 75 76 77 78 79 80 81 82 83 84 85 86 87 88 89 90 91 92 93 94 95 96 97 98 99 100 101 102 103 104 105 106 107 108 109 110 111 112 113 114 115 116 117 118 119 120 121 122 123 124 125 126 127 128 129 130 131 132 133 134 135 136 137 138 139 140 141 142 143 144 145 146 147 148 149 150 151 152 153 154 155 156 157 158 159 160 161 162 163 164 165 166 167 168 169 170 171 172 173 174 175 176 177 178 179 180 181 182 183 184 185 186 187 188 189 190 191 192 193 194 195 196 197 198 199 200 201 202 203 204 205 206 207 208 209 210 211 212 213 214 215 216 217 218 219 220 221 222 223 224 225 226 227 228 229 230 231 232 233 234 235 236 237 238 239 240 241 242 243 244 245 246 247 248 249 250 251 252 253 254 255 256 257 258 259 260 261 262 263 264 265 266 267 268 269 270 271 272 273 274 275 276 277 278 279 280 281 282 283 284 285 286 287 288 289 290 291 292 293 294 295 296 297 298 299 300 301 302 303 304 305 306 307 308 309 310 311 312 313 314 315 316 317 318 319 320 321 322 323 324 325 326 327 328 329 330 331 332 333 334 335 336 337 338 339 340 341 342 343 344 345 346 347 348 349 350 351 352 353 354 355 356 357 358 359 360 361 362 363 364 365 366 367 368 369 370 371 372 373 374 375 376 377 378 379 380 381 382 383 384 385 386 387 388 389 390 391 392 393 394 395 396 397 398 399 400 401 402 403 404 405 406 407 408 409 410 411 412 413 414 415 416 417 418 419 420 421 422 423 424 425 426 427 428 429 430 431 432 433 434 435 436 437 438 439 440 441 442 443 444 445 446 447 448 449 450 451 452 453 454 455 456 457 458 459 460 461 462 463 464 465 466 467 468 469 470 471 472 473 474 475 476 477 478 479 480 481 482 483 484 485 486 487 488 489 490 491 492 493 494 495 496 497 498 499 500 501 502 503 504 505 506 507 508 509 510 511 512 513 514 515 516 517 518 519 520 521 522 523 524 525 526 527 528 529 530 531 532 533 534 535 536 537 538 539 540 541 542 543 544 545 546 547 548 549 550 551 552 553 554 555 556 557 558 559 560 561 562 563 564 565 566 567 568 569 570 571 572 573 574 575 576 577 578 579 580 581 582 583 584 585 586 587 588 589 590 591 592 593 594 595 596 597 598 599 600 601 602 603 604 605 606 607 608 609 610 611 612 613 614 615 616 617 618 619 620 621 622 623 624 625 626 627 628 629 630 631 632 633 634 635 636 637 638 639 640 641 642 643 644 645 646 647 648 649 650 651 652 653 654 655 656 657 658 659 660 661 662 663 664 665 666 667 668 669 670 671 672 673 674 675 676 677 678 679 680 681 682 683 684 685 686 687 688 689 690 691 692 693 694 695 696 697 698 699 700 701 702 703 704 705 706 707 708 709 710 711 712 713 714 715 716 717 718 719 720 721 722 723 724 725 726 727 728 729 730 731 732 733 734 735 736 737 738 739 740 741 742 743 744 745 746 747 748 749 750 751 752 753 754 755 756 757 758 759 760 761 762 763 764 765 766 767 768 769 770 771 772 773 774 775 776 777 778 779 780 781 782 783 784 785 786 787 788 789 790 791 792 793 794 795 796 797 798 799 800 801 802 803 804 805 806 807 808 809 810 811 812 813 814 815 816 817 818 819 820 821 822 823 824 825 826 827 828 829 830 831 832 833 834 835 836 837 838 839 840 841 842 843 844 845 846 847 848 849 850 851 852 853 854 855 856 857 858 859 860 861 862 863 864 865 866 867 868 869 870 871 872 873 874 875 876 877 878 879 880 881 882 883 884 885 886 887 888 889 890 891 892 893 894 895 896 897 898 899 900 901 902 903 904 905 906 907 908 909 910 911 912 913 914 915 916 917 918 919 920 921 922 923 924 925 926 927 928 929 930 931 932 933 934 935 936 937 938 939 940 941 942 943 944 945 946 947 948 949 950 951 952 953 954 955 956 957 958 959 960 961 962 963 964 965 966 967 968 969 970 971 972 973 974 975 976 977 978 979 980 981 982 983 984 985 986 987 988 989 990 991 992 993 994 995 996 997 998 999 1000

Leu Glu Gln Ser Leu Thr Phe Val Gln Ala Leu Ile Glu Lys Asp Ala  
 115 120 125  
 Ser Asp Val Lys Cys Ala Asn Gly Asp Pro Pro Ile Phe Thr Lys Pro  
 130 135 140  
 Asp Lys Ile Ser Gly Val Ile Gly Ala Ala Ala Ser Ser Val Ser Ile  
 145 150 155 160  
 Met Val Ala Asn Ile Leu Arg Leu Phe Lys Ile Pro Gln Ile Ser Tyr  
 165 170 175  
 Ala Ser Thr Ala Pro Glu Leu Ser Asp Asn Thr Arg Tyr Asp Phe Phe  
 180 185 190  
 Ser Arg Val Val Pro Pro Asp Ser Tyr Gln Ala Gln Ala Met Val Asp  
 195 200 205  
 Ile Val Thr Ala Leu Gly Trp Asn Tyr Val Ser Thr Leu Ala Ser Glu  
 210 215 220  
 Gly Asn Tyr Gly Glu Ser Gly Val Glu Ala Phe Thr Gln Ile Ser Arg  
 225 230 235 240  
 Glu Ile Gly Gly Val Cys Ile Ala Gln Ser Gln Lys Ile Pro Arg Glu  
 245 250 255  
 Pro Arg Pro Gly Glu Phe Glu Lys Ile Ile Lys Arg Leu Leu Glu Thr  
 260 265 270  
 Pro Asn Ala Arg Ala Val Ile Met Phe Ala Asn Glu Asp Asp Ile Arg  
 275 280 285  
 Arg Ile Leu Glu Ala Ala Lys Lys Leu Asn Gln Ser Gly His Phe Leu  
 290 295 300  
 Trp Ile Gly Ser Asp Ser Trp Gly Ser Lys Ile Ala Pro Val Tyr Gln  
 305 310 315 320  
 Gln Glu Glu Ile Ala Glu Gly Ala Val Thr Ile Leu Pro Lys Arg Ala  
 325 330 335  
 Ser Ile Asp Gly Phe Asp Arg Tyr Phe Arg Ser Arg Thr Leu Ala Asn  
 340 345 350  
 Asn Arg Arg Asn Val Trp Phe Ala Glu Phe Trp Glu Glu Asn Phe Gly  
 355 360 365  
 Cys Lys Leu Gly Ser His Gly Lys Arg Asn Ser His Ile Lys Lys Cys  
 370 375 380  
 Thr Gly Leu Glu Arg Ile Ala Arg Asp Ser Ser Tyr Glu Gln Glu Gly  
 385 390 395 400  
 Lys Val Gln Phe Val Ile Asp Ala Val Tyr Ser Met Ala Tyr Ala Leu  
 405 410 415

His Asn Met His Lys Asp Leu Cys Pro Gly Tyr Ile Gly Leu Cys Pro  
 420 425 430  
 Arg Met Ser Thr Ile Asp Gly Lys Glu Leu Leu Gly Tyr Ile Arg Ala  
 435 440 445  
 Val Asn Phe Asn Gly Ser Ala Gly Thr Pro Val Thr Phe Asn Glu Asn  
 450 455 460  
 Gly Asp Ala Pro Gly Arg Tyr Asp Ile Phe Gln Tyr Gln Ile Thr Asn  
 465 470 475 480  
 Lys Ser Thr Glu Tyr Lys Val Ile Gly His Trp Thr Asn Gln Leu His  
 485 490 495  
 Leu Lys Val Glu Asp Met Gln Trp Ala His Arg Glu His Thr His Pro  
 500 505 510  
 Ala Ser Val Cys Ser Leu Pro Cys Lys Pro Gly Glu Arg Lys Lys Thr  
 515 520 525  
 Val Lys Gly Val Pro Cys Cys Trp His Cys Glu Arg Cys Glu Gly Tyr  
 530 535 540  
 Asn Tyr Gln Val Asp Glu Leu Ser Cys Glu Leu Cys Pro Leu Asp Gln  
 545 550 555 560  
 Arg Pro Asn Met Asn Arg Thr Gly Cys Gln Leu Ile Pro Ile Ile Lys  
 565 570 575  
 Leu Glu Trp His Ser Pro Trp  
 580

<210> 6  
 <211> 250  
 <212> PRT  
 <213> Human

<400> 6

Gly Ile Ala Leu Thr Leu Phe Ala Val Leu Gly Ile Phe Leu Thr Ala  
 1 5 10 15  
 Phe Val Leu Gly Val Phe Ile Lys Phe Arg Asn Thr Pro Ile Val Lys  
 20 25 30  
 Ala Thr Asn Arg Glu Leu Ser Tyr Leu Leu Leu Phe Ser Leu Leu Cys  
 35 40 45  
 Cys Phe Ser Ser Ser Leu Phe Phe Ile Gly Glu Pro Gln Asp Trp Thr  
 50 55 60  
 Cys Arg Leu Arg Gln Pro Ala Phe Gly Ile Ser Phe Val Leu Cys Ile



65	70	75	80
Ser Cys Ile Leu Val Lys Thr Asn Arg Val Leu Leu Val Phe Glu Ala	85	90	95
Lys Ile Pro Thr Ser Phe His Arg Lys Trp Trp Gly Leu Asn Leu Gln	100	105	110
Phe Leu Leu Val Phe Leu Cys Thr Phe Met Gln Ile Val Ile Cys Val	115	120	125
Ile Trp Leu Tyr Thr Ala Pro Pro Ser Ser Tyr Arg Asn Gln Glu Leu	130	135	140
Glu Asp Glu Ile Ile Phe Ile Thr Cys His Glu Gly Ser Leu Met Ala	145	150	155
Leu Gly Phe Leu Ile Gly Tyr Thr Cys Leu Leu Ala Ala Ile Cys Phe	165	170	175
Phe Phe Ala Phe Lys Ser Arg Lys Leu Pro Glu Asn Phe Asn Glu Ala	180	185	190
Lys Phe Ile Thr Phe Ser Met Leu Ile Phe Phe Ile Val Trp Ile Ser	195	200	205
Phe Ile Pro Ala Tyr Ala Ser Thr Tyr Gly Lys Phe Val Ser Ala Val	210	215	220
Glu Val Ile Ala Ile Leu Ala Ala Ser Phe Gly Leu Leu Ala Cys Ile	225	230	235
Phe Phe Asn Lys Ile Tyr Ile Ile Leu Phe	245	250	

<210> 7  
 <211> 267  
 <212> PRT  
 <213> Human

<400> 7

Leu Phe Ile Ser Val Ser Val Leu Ser Ser Leu Gly Ile Val Leu Ala	1	5	10	15
Val Val Cys Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile	20	25	30	
Gln Asn Ser Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser	35	40	45	
Leu Ala Leu Ala Ala Val Phe Pro Leu Gly Leu Asp Gly Tyr His Ile	50	55	60	

Gly Arg Asn Gln Phe Pro Phe Val Cys Gln Ala Arg Leu Trp Leu Leu  
 65 70 75 80  
 Gly Leu Gly Phe Ser Leu Gly Tyr Gly Ser Met Phe Thr Lys Ile Trp  
 85 90 95  
 Trp Val His Thr Val Phe Thr Lys Lys Glu Glu Lys Lys Glu Trp Arg  
 100 105 110  
 Lys Thr Leu Glu Pro Trp Lys Leu Tyr Ala Thr Val Gly Leu Leu Val  
 115 120 125  
 Gly Met Asp Val Leu Thr Leu Ala Ile Trp Gln Ile Val Asp Pro Leu  
 130 135 140  
 His Arg Thr Ile Glu Thr Phe Ala Lys Glu Glu Pro Lys Glu Asp Ile  
 145 150 155 160  
 Asp Val Ser Ile Leu Pro Gln Leu Glu His Cys Ser Ser Arg Lys Met  
 165 170 175  
 Asn Thr Trp Leu Gly Ile Phe Tyr Gly Tyr Lys Gly Leu Leu Leu Leu  
 180 185 190  
 Leu Gly Ile Phe Leu Ala Tyr Glu Thr Lys Ser Val Ser Thr Glu Lys  
 195 200 205  
 Ile Asn Asp His Arg Ala Val Gly Met Ala Ile Tyr Asn Val Ala Val  
 210 215 220  
 Leu Cys Leu Ile Thr Ala Pro Val Thr Met Ile Leu Ser Ser Gln Gln  
 225 230 235 240  
 Asp Ala Ala Phe Ala Phe Ala Ser Leu Ala Ile Val Phe Ser Ser Tyr  
 245 250 255  
 Ile Thr Leu Val Val Leu Phe Val Pro Lys Met  
 260 265

<210> 8  
 <211> 267  
 <212> PRT  
 <213> Human

<400> 8

Leu Phe Ile Ser Val Ser Val Leu Ser Ser Leu Gly Ile Val Leu Ala  
 1 5 10 15  
 Val Val Cys Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile  
 20 25 30  
 Gln Asn Ser Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser  
 35 40 45

Leu Ala Leu Ala Ala Val Phe Pro Leu Gly Leu Asp Gly Tyr His Ile  
 50 55 60  
 Gly Arg Asn Gln Phe Pro Phe Val Cys Gln Ala Arg Leu Trp Leu Leu  
 65 70 75 80  
 Gly Leu Gly Phe Ser Leu Gly Tyr Gly Ser Met Phe Thr Lys Ile Trp  
 85 90 95  
 Trp Val His Thr Val Phe Thr Lys Lys Glu Glu Lys Lys Glu Trp Arg  
 100 105 110  
 Lys Thr Leu Glu Pro Trp Lys Leu Tyr Ala Thr Val Gly Leu Leu Val  
 115 120 125  
 Gly Met Asp Val Leu Thr Leu Ala Ile Trp Gln Ile Val Asp Pro Leu  
 130 135 140  
 His Arg Thr Ile Glu Thr Phe Ala Lys Glu Glu Pro Lys Glu Asp Ile  
 145 150 155 160  
 Asp Val Ser Ile Leu Pro Gln Leu Glu His Cys Ser Ser Arg Lys Met  
 165 170 175  
 Asn Thr Trp Leu Gly Ile Phe Tyr Gly Tyr Lys Gly Leu Leu Leu Leu  
 180 185 190  
 Leu Gly Ile Phe Leu Ala Tyr Glu Thr Lys Ser Val Ser Thr Glu Lys  
 195 200 205  
 Ile Asn Asp His Arg Ala Val Gly Met Ala Ile Tyr Asn Val Ala Val  
 210 215 220  
 Leu Cys Leu Ile Thr Ala Pro Val Thr Met Ile Leu Ser Ser Gln Gln  
 225 230 235 240  
 Asp Ala Ala Phe Ala Phe Ala Ser Leu Ala Ile Val Phe Ser Ser Tyr  
 245 250 255  
 Ile Thr Leu Val Val Leu Phe Val Pro Lys Met  
 260 265

<210> 9  
 <211> 264  
 <212> PRT  
 <213> Human

<400> 9

Leu Tyr Ser Ile Leu Ser Ala Leu Thr Ile Leu Gly Met Ile Met Ala  
 1 5 10 15

Ser Ala Phe Leu Phe Phe Asn Ile Lys Asn Arg Asn Gln Lys Leu Ile  
 20 25 30  
 Lys Met Ser Ser Pro Tyr Met Asn Asn Leu Ile Ile Leu Gly Gly Met  
 35 40 45  
 Leu Ser Tyr Ala Ser Ile Phe Leu Phe Gly Leu Asp Gly Ser Phe Val  
 50 55 60  
 Ser Glu Lys Thr Phe Glu Thr Leu Cys Thr Val Arg Thr Trp Ile Leu  
 65 70 75 80  
 Thr Val Gly Tyr Thr Thr Ala Phe Gly Ala Met Phe Ala Lys Thr Trp  
 85 90 95  
 Arg Val His Ala Ile Phe Lys Asn Val Lys Met Lys Lys Lys Ile Ile  
 100 105 110  
 Lys Asp Gln Lys Leu Leu Val Ile Val Gly Gly Met Leu Leu Ile Asp  
 115 120 125  
 Leu Cys Ile Leu Ile Cys Trp Gln Ala Val Asp Pro Leu Arg Arg Thr  
 130 135 140  
 Val Glu Lys Tyr Ser Met Glu Pro Asp Pro Ala Gly Arg Asp Ile Ser  
 145 150 155 160  
 Ile Arg Pro Leu Leu Glu His Cys Glu Asn Thr His Met Thr Ile Trp  
 165 170 175  
 Leu Gly Ile Val Tyr Ala Tyr Lys Gly Leu Leu Met Leu Phe Gly Cys  
 180 185 190  
 Phe Leu Ala Trp Glu Thr Arg Asn Val Ser Ile Pro Ala Leu Asn Asp  
 195 200 205  
 Ser Lys Tyr Ile Gly Met Ser Val Tyr Asn Val Gly Ile Met Cys Ile  
 210 215 220  
 Ile Gly Ala Ala Val Ser Phe Leu Thr Arg Asp Gln Pro Asn Val Gln  
 225 230 235 240  
 Phe Cys Ile Val Ala Leu Val Ile Ile Phe Cys Ser Thr Ile Thr Leu  
 245 250 255  
 Cys Leu Val Phe Val Pro Lys Leu  
 260

[illegible]

<210> 11  
 <211> 216  
 <212> PRT  
 <213> Human

<400> 11

```

Lys Pro Ser Arg Asn Thr Ile Glu Glu Val Arg Cys Ser Thr Ala Ala
 1          5          10          15

His Ala Phe Lys Val Ala Ala Arg Ala Thr Leu Arg Arg Ser Asn Val
      20          25          30

Ser Arg Lys Arg Ser Ser Ser Leu Gly Gly Ser Thr Gly Ser Thr Pro
      35          40          45

Ser Ser Ser Ile Ser Ser Lys Ser Asn Ser Glu Asp Pro Phe Pro Gln
 50          55          60

Pro Glu Arg Gln Lys Gln Gln Gln Pro Leu Ala Leu Thr Gln Gln Glu
 65          70          75          80

Gln Gln Gln Gln Pro Leu Thr Leu Pro Gln Gln Gln Arg Ser Gln Gln
      85          90          95

Gln Pro Arg Cys Lys Gln Lys Val Ile Phe Gly Ser Gly Thr Val Thr
      100          105          110

Phe Ser Leu Ser Phe Asp Glu Pro Gln Lys Asn Ala Met Ala His Gly
      115          120          125

Asn Ser Thr His Gln Asn Ser Leu Glu Ala Gln Lys Ser Ser Asp Thr
      130          135          140

Leu Thr Arg His Gln Pro Leu Leu Pro Leu Gln Cys Gly Glu Thr Asp
      145          150          155          160

Leu Asp Leu Thr Val Gln Glu Thr Gly Leu Gln Gly Pro Val Gly Gly
      165          170          175

Asp Gln Arg Pro Glu Val Glu Asp Pro Glu Glu Leu Ser Pro Ala Leu
      180          185          190

Val Val Ser Ser Ser Gln Ser Phe Val Ile Ser Gly Gly Gly Ser Thr
      195          200          205

Val Thr Glu Asn Val Val Asn Ser
      210          215

```

<210> 12  
 <211> 104  
 <212> PRT  
 <213> Human

<400> 12

```

Arg Arg Leu Ile Thr Arg Gly Glu Trp Gln Ser Glu Ala Gln Asp Thr
 1           5           10           15

Met Lys Thr Gly Ser Ser Thr Asn Asn Asn Glu Glu Glu Lys Ser Arg
          20           25           30

Leu Leu Glu Lys Glu Asn Arg Glu Leu Glu Lys Ile Ile Ala Glu Lys
      35           40           45

Glu Glu Arg Val Ser Glu Leu Arg His Gln Leu Gln Ser Arg Gln Gln
 50           55           60

Leu Arg Ser Arg Arg His Pro Pro Thr Pro Pro Glu Pro Ser Gly Gly
65           70           75           80

Leu Pro Arg Gly Pro Pro Glu Pro Pro Asp Arg Leu Ser Cys Asp Gly
          85           90           95

Ser Arg Val His Leu Leu Tyr Lys
          100
  
```

<210> 13  
 <211> 104  
 <212> PRT  
 <213> Human

<400> 13

```

Arg Arg Leu Ile Thr Arg Gly Glu Trp Gln Ser Glu Ala Gln Asp Thr
 1           5           10           15

Met Lys Thr Gly Ser Ser Thr Asn Asn Asn Glu Glu Glu Lys Ser Arg
          20           25           30

Leu Leu Glu Lys Glu Asn Arg Glu Leu Glu Lys Ile Ile Ala Glu Lys
      35           40           45

Glu Glu Arg Val Ser Glu Leu Arg His Gln Leu Gln Ser Arg Gln Gln
 50           55           60

Leu Arg Ser Arg Arg His Pro Pro Thr Pro Pro Glu Pro Ser Gly Gly
65           70           75           80

Leu Pro Arg Gly Pro Pro Glu Pro Pro Asp Arg Leu Ser Cys Asp Gly
          85           90           95

Ser Arg Val His Leu Leu Tyr Lys
          100
  
```

<210> 14  
 <211> 197  
 <212> PRT  
 <213> Human

<400> 14

```

Ile Thr Leu Arg Thr Asn Pro Asp Ala Ala Thr Gln Asn Arg Arg Phe
 1           5           10           15

Gln Phe Thr Gln Asn Gln Lys Lys Glu Asp Ser Lys Thr Ser Thr Ser
      20           25           30

Val Thr Ser Val Asn Gln Ala Ser Thr Ser Arg Leu Glu Gly Leu Gln
      35           40           45

Ser Glu Asn His Arg Leu Arg Met Lys Ile Thr Glu Leu Asp Lys Asp
 50           55           60

Leu Glu Glu Val Thr Met Gln Leu Gln Asp Thr Pro Glu Lys Thr Thr
 65           70           75           80

Tyr Ile Lys Gln Asn His Tyr Gln Glu Leu Asn Asp Ile Leu Asn Leu
      85           90           95

Gly Asn Phe Thr Glu Ser Thr Asp Gly Gly Lys Ala Ile Leu Lys Asn
      100          105          110

His Leu Asp Gln Asn Pro Gln Leu Gln Trp Asn Thr Thr Glu Pro Ser
      115          120          125

Arg Thr Cys Lys Asp Pro Ile Glu Asp Ile Asn Ser Pro Glu His Ile
      130          135          140

Gln Arg Arg Leu Ser Leu Gln Leu Pro Ile Leu His His Ala Tyr Leu
      145          150          155          160

Pro Ser Ile Gly Gly Val Asp Ala Ser Cys Val Ser Pro Cys Val Ser
      165          170          175

Pro Thr Ala Ser Pro Arg His Arg His Val Pro Pro Ser Phe Arg Val
      180          185          190

Met Val Ser Gly Leu
      195

```



<210> 15  
 <211> 65  
 <212> PRT  
 <213> Human

<400> 15

```

His Pro Glu Gln Asn Val Gln Lys Arg Lys Arg Ser Phe Lys Ala Val
 1          5          10          15
Val Thr Ala Ala Thr Met Gln Ser Lys Leu Ile Gln Lys Gly Asn Asp
          20          25          30
Arg Pro Asn Gly Glu Val Lys Ser Glu Leu Cys Glu Ser Leu Glu Thr
          35          40          45
Asn Ser Lys Ser Ser Val Glu Phe Pro Met Val Lys Ser Gly Ser Thr
 50          55          60
Ser
65

```

<210> 16  
 <211> 374  
 <212> PRT  
 <213> Human

<400> 16

```

Met Ala Arg Ser Leu Thr Trp Gly Cys Cys Pro Trp Cys Leu Thr Glu
 1          5          10          15
Glu Glu Lys Thr Ala Ala Arg Ile Asp Gln Glu Ile Asn Arg Ile Leu
          20          25          30
Leu Glu Gln Lys Lys Gln Glu Arg Glu Glu Leu Lys Leu Leu Leu Leu
          35          40          45
Gly Pro Gly Glu Ser Gly Lys Ser Thr Phe Ile Lys Gln Met Arg Ile
 50          55          60
Ile His Gly Val Gly Tyr Ser Glu Glu Asp Arg Arg Ala Phe Arg Leu
65          70          75          80
Leu Ile Tyr Gln Asn Ile Phe Val Ser Met Gln Ala Met Ile Asp Ala
          85          90          95
Met Asp Arg Leu Gln Ile Pro Phe Ser Arg Pro Asp Ser Lys Gln His
          100          105          110
Ala Ser Leu Val Met Thr Gln Asp Pro Tyr Lys Val Ser Thr Phe Glu
          115          120          125
Lys Pro Tyr Ala Val Ala Met Gln Tyr Leu Trp Arg Asp Ala Gly Ile
          130          135          140

```

Arg Ala Cys Tyr Glu Arg Arg Arg Glu Phe His Leu Leu Asp Ser Ala  
 145 150 155 160  
 Val Tyr Tyr Leu Ser His Leu Glu Arg Ile Ser Glu Asp Ser Tyr Ile  
 165 170 175  
 Pro Thr Ala Gln Asp Val Leu Arg Ser Arg Met Pro Thr Thr Gly Ile  
 180 185 190  
 Asn Glu Tyr Cys Phe Ser Val Lys Lys Thr Lys Leu Arg Ile Val Asp  
 195 200 205  
 Val Gly Gly Gln Arg Ser Glu Arg Arg Lys Trp Ile His Cys Phe Glu  
 210 215 220  
 Asn Val Ile Ala Leu Ile Tyr Leu Ala Ser Leu Ser Glu Tyr Asp Gln  
 225 230 235 240  
 Cys Leu Glu Glu Asn Asp Gln Glu Asn Arg Met Glu Glu Ser Leu Ala  
 245 250 255  
 Leu Phe Ser Thr Ile Leu Glu Leu Pro Trp Phe Lys Ser Thr Ser Val  
 260 265 270  
 Ile Leu Phe Leu Asn Lys Thr Asp Ile Leu Glu Asp Lys Ile His Thr  
 275 280 285  
 Ser His Leu Ala Thr Tyr Phe Pro Ser Phe Gln Gly Pro Arg Arg Asp  
 290 295 300  
 Ala Glu Ala Ala Lys Ser Phe Ile Leu Asp Met Tyr Ala Arg Val Tyr  
 305 310 315 320  
 Ala Ser Cys Ala Glu Pro Gln Asp Gly Gly Arg Lys Gly Ser Arg Ala  
 325 330 335  
 Arg Arg Phe Phe Ala His Phe Thr Cys Ala Thr Asp Thr Gln Ser Val  
 340 345 350  
 Arg Ser Val Phe Lys Asp Val Arg Asp Ser Val Leu Ala Arg Tyr Leu  
 355 360 365  
 Asp Glu Ile Asn Leu Leu  
 370

<210> 17  
 <211> 374  
 <212> PRT  
 <213> Human

<400> 17

```

Met Ala Arg Ser Leu Thr Trp Arg Cys Cys Pro Trp Cys Leu Thr Glu
 1          5          10          15

Asp Glu Lys Ala Ala Ala Arg Val Asp Gln Glu Ile Asn Arg Ile Leu
 20          25          30

Leu Glu Gln Lys Lys Gln Asp Arg Gly Glu Leu Lys Leu Leu Leu Leu
 35          40          45

Gly Pro Gly Glu Ser Gly Lys Ser Thr Phe Ile Lys Gln Met Arg Ile
 50          55          60

Ile His Gly Ala Gly Tyr Ser Glu Glu Glu Arg Lys Gly Phe Arg Pro
 65          70          75          80

Leu Val Tyr Gln Asn Ile Phe Val Ser Met Arg Ala Met Ile Glu Ala
          85          90          95

Met Glu Arg Leu Gln Ile Pro Phe Ser Arg Pro Glu Ser Lys His His
          100          105          110

Ala Ser Leu Val Met Ser Gln Asp Pro Tyr Lys Val Thr Thr Phe Glu
          115          120          125

Lys Arg Tyr Ala Ala Ala Met Gln Trp Leu Trp Arg Asp Ala Gly Ile
          130          135          140

Arg Ala Cys Tyr Glu Arg Arg Arg Glu Phe His Leu Leu Asp Ser Ala
          145          150          155          160

Val Tyr Tyr Leu Ser His Leu Glu Arg Ile Thr Glu Glu Gly Tyr Val
          165          170          175

Pro Thr Ala Gln Asp Val Leu Arg Ser Arg Met Pro Thr Thr Gly Ile
          180          185          190

Asn Glu Tyr Cys Phe Ser Val Gln Lys Thr Asn Leu Arg Ile Val Asp
          195          200          205

Val Gly Gly Gln Lys Ser Glu Arg Lys Lys Trp Ile His Cys Phe Glu
          210          215          220

Asn Val Ile Ala Leu Ile Tyr Leu Ala Ser Leu Ser Glu Tyr Asp Gln
          225          230          235          240

Cys Leu Glu Glu Asn Asn Gln Glu Asn Arg Met Lys Glu Ser Leu Ala
          245          250          255

Leu Phe Gly Thr Ile Leu Glu Leu Pro Trp Phe Lys Ser Thr Ser Val
          260          265          270

```

Ile Leu Phe Leu Asn Lys Thr Asp Ile Leu Glu Glu Lys Ile Pro Thr  
 275 280 285

Ser His Leu Ala Thr Tyr Phe Pro Ser Phe Gln Gly Pro Lys Gln Asp  
 290 295 300

Ala Glu Ala Ala Lys Arg Phe Ile Leu Asp Met Tyr Thr Arg Met Tyr  
 305 310 315 320

Thr Gly Cys Val Asp Gly Pro Glu Gly Ser Lys Lys Gly Ala Arg Ser  
 325 330 335

Arg Arg Leu Phe Ser His Tyr Thr Cys Ala Thr Asp Thr Gln Asn Ile  
 340 345 350

Arg Lys Val Phe Lys Asp Val Arg Asp Ser Val Leu Ala Arg Tyr Leu  
 355 360 365

Asp Glu Ile Asn Leu Leu  
 370

<210> 18  
 <211> 3234  
 <212> DNA  
 <213> Human

<400> 18

atggcatttt	atagctgctg	ctgggtcctc	ttggcactca	cctggcacac	ctctgcctac	60
gggccagacc	agcgagccca	aaagaagggg	gacattatcc	ttgggggggt	ctttcctatt	120
cattttggag	tagcagctaa	agatcaagat	ctcaaataca	ggccggagtc	tgtggaatgt	180
atcaggtata	atttccgtgg	gtttcgctgg	ttacaggcta	tgatatttgc	catagaggag	240
ataaacagca	gcccagccct	tcttcccaac	ttgacgctgg	gatacaggat	atttgacact	300
tgcaacaccg	tttctaaggg	cttgggaagcc	accctgagtt	ttgttgctca	aaacaaaatt	360
gattccttga	accttgatga	gttctgcaac	tgctcagagc	acattccctc	tacgattgct	420
gtggtgggag	caactggctc	aggcgtctcc	acggcagtg	caaactctgc	ggggctcttc	480
tacattcccc	aggtcagtta	tgctctctcc	agcagactcc	tcagcaacaa	gaatcaattc	540
aagtctttcc	tccgaaccat	ccccaatgat	gagcaccagg	ccactgccat	ggcagacatc	600
atcgagtatt	tccgctggaa	ctgggtgggc	acaattgcag	ctgatgacga	ctatgggcgg	660
ccggggattg	agaaattccg	agaggaagct	gaggaaaggg	atatctgcat	cgacttcagt	720
gaactcatct	cccagtactc	tgatgaggaa	gagatccagc	atgtggtaga	ggtgattcaa	780
aattccacgg	ccaaagtcac	cgtggttttc	tccagtggcc	cagatcttga	gcccctcatc	840
aaggagattg	tccggcgcaa	tatcacgggc	aagatctggc	tggccagcga	ggcctggggc	900
agctcctccc	tgatcgccat	gcctcagtac	ttccacgtgg	ttggcggcac	cattggattc	960
gctctgaagg	ctgggcagat	cccaggcttc	cgggaattcc	tgaagaaggt	ccatcccagg	1020
aagtctgtcc	acaatggttt	tgccaaggag	ttttgggaag	aaacatttaa	ctgccacctc	1080
caagaagggtg	caaaaggacc	tttacctgtg	gacacctttc	tgagaggtca	cgaagaaagt	1140
ggcgacaggt	ttagcaacag	ctcgacagcc	ttccgacccc	tctgtacagg	ggatgagaac	1200
atcagcagtg	tcgagacccc	ttacatagat	tacacgcatt	tacggatata	ctacaatgtg	1260
tacttagcag	tctactccat	tgcccacgcc	ttgcaagata	tatatacctg	cttacctggg	1320
agagggctct	tcaccaatgg	ctcctgtgca	gacatcaaga	aagttgaggc	gtggcagggtc	1380
ctgaagcacc	tacggcatct	aaactttaca	aacaatatgg	gggagcaggt	gacctttgat	1440
gagtgtggtg	acctggtggg	gaactattcc	atcatcaact	ggcacctctc	cccagaggat	1500

ggctccatcg	tgtttaagga	agtcgggtat	tacaacgtct	atgccaagaa	gggagaaaga	1560
ctcttcatca	acgaggagaa	aatcctgtgg	agtgggttct	ccagggaggt	gcccttctcc	1620
aactgcagcc	gagactgcct	ggcagggacc	aggaaaggga	tcattgaggg	ggagcccacc	1680
tgttgctttg	agtgtgtgga	gtgtcctgat	ggggagtata	gtgatgagac	agatgccagt	1740
gcctgtaaca	agtgccaga	tgacttcttg	tccaatgaga	accacacctc	ctgcattgcc	1800
aaggagatcg	agtttctgtc	gtggacggag	ccctttggga	tcgcactcac	cctctttgcc	1860
gtgctgggca	ttttcctgac	agcctttgtg	ctgggtgtgt	ttatcaagtt	ccgcaacaca	1920
cccattgtca	aggccacca	ccgagagctc	tcctacctcc	tcctcttctc	cctgctctgc	1980
tgettctcca	gctccctggt	cttcatcggg	gagccccagg	actggacgtg	ccgcctgcgc	2040
cagccggcct	ttggcatcag	cttcgtgtct	tgcattctcat	gcattcctgg	gaaaaccaac	2100
cgtgtcctcc	tgggtgttga	ggccaagatc	cccaccagct	tccaccgcaa	gtggtggggg	2160
ctcaacctgc	agttcctgct	ggttttcctc	tgcaccttca	tgcagattgt	catctgtgtg	2220
atctggctct	acaccgcgcc	ccctcacaag	taccgcaacc	aggagctgga	ggatgagatc	2280
atcttcatca	cgtgccacga	gggctccctc	atggccctgg	gcttcctgat	cggctacacc	2340
tgctgctgg	ctgccatctg	cttcttcttt	gccttcaagt	cccggaagct	gccggagaac	2400
ttcaatgaag	ccaagtcat	caccttcagc	atgctcatct	tcttcatcgt	ctggatctcc	2460
ttcattccag	cctatgccag	cacctatggc	aagtttgtct	ctgccgtaga	ggtgattgcc	2520
atcctggcag	ccagctttgg	cttgctggcg	tgcattctct	tcaacaagat	ctacatcatt	2580
ctcttcaagc	catcccgcga	caccatcgag	gaggtgcgtt	gcagcaccgc	agctcacgct	2640
ttcaaggtgg	ctgcccgggc	cacgctgcgc	cgcagcaacg	tctcccgcga	gcggtccagc	2700
agccttggag	gctccacggg	atccaccccc	tcctcctcca	tcagcagcaa	gagcaacagc	2760
gaagacccat	tcacacagcc	cgagaggcag	aagcagcagc	agccgctggc	cctaaccagc	2820
caagagcagc	agcagcagcc	cctgaccctc	ccacagcagc	aacgatctca	gcagcagccc	2880
agatgcaagc	agaaggtcat	ctttggcagc	ggcacgggtc	ccttctcact	gagctttgat	2940
gagcctcaga	agaacgccat	ggcccacggg	aattctacgc	accagaactc	cctggaggcc	3000
cagaaaagca	gcgatacgct	gacccgacac	cagccattac	tcccgtgca	gtgcggggaa	3060
acggacttag	atctgaccgt	ccaggaaaaca	ggtctgcaag	gacctgtggg	tggagaccag	3120
cggccagagg	tggaggaccc	tgaagagttg	tcccagcac	ttgtagtgtc	cagttcacag	3180
agctttgtca	tcagtgggtg	aggcagcact	gttacagaaa	acgtagtga	ttca	3234

<210> 19  
 <211> 3464  
 <212> DNA  
 <213> Human

<400> 19

atgttgetgc	tgetgctact	ggcgccactc	ttcctccgcc	ccccgggcgc	gggcggggcg	60
cagacccccca	acgccacctc	agaaggttgc	cagatcatac	accgcacctg	ggaagggggc	120
atcaggtacc	ggggcctgac	tccggaccag	gtgaaggcta	tcaacttctc	gccagtggac	180
tatgagattg	agtatgtgtg	ccggggggag	cgcgaggtgg	tggggcccaa	ggtccgcaag	240
tgctgggcca	acggctcctg	gacagatatg	gacacaccca	gccgctgtgt	ccgaatctgc	300
tccaagtctt	atttgacctt	ggaaaatggg	aaggttttcc	tgacgggtgg	ggacctccca	360
gctctggacg	gagcccgggt	ggatttccgg	tgtgaccccc	acttccatct	ggtgggcagc	420
tcccggagca	tctgtagtca	gggcccagtgg	agcaccacca	agccccactg	ccaggtgaat	480
cgaacgccac	actcagaacg	gcgcgcagtg	tacatcgggg	cactgtttcc	catgagcggg	540
ggctggccag	ggggccaggc	ctgccagccc	gcgggtggaga	tggcgtgga	ggacgtgaat	600
agccgcaggg	acatcctgcc	ggactatgag	ctcaagctca	tccaccacga	cagcaagtgt	660
gatccaggcc	aagccacca	gtacctatat	gagctgctct	acaacgaccc	tatcaagatc	720
atccttatgc	ctggctgcag	ctctgtctcc	acgctgggtg	ctgaggctgc	taggatgtgg	780
aacctcattg	tgttttctta	tggctccagc	tcaccagccc	tgtcaaaccg	gcagcgtttc	840
cccactttct	tccgaacgca	cccatcagcc	acactccaca	accctaccgc	cgtgaaactc	900
tttgaaaagt	ggggctggaa	gaagattgct	accatccagc	agaccactga	ggtcttctct	960
tcgactctgg	acgacctgga	ggaacgagtg	aaggaggctg	gaattgagat	tactttccgc	1020

cagagtttct	tctcagatcc	agctgtgccc	gtcaaaaacc	tgaagcgcca	ggatgcccga	1080
atcatcgtgg	gactttttcta	tgagactgaa	gcccggaaag	ttttttgtga	ggtgtacaag	1140
gagcgtctct	ttgggaagaa	gtacgtctgg	ttcctcattg	ggtggtatgc	tgacaattgg	1200
ttcaagatct	acgaccttcc	tatcaactgc	acagtggatg	agatgactga	ggcggtagag	1260
ggccacatca	caactgagat	tgtcatgctg	aatcctgcca	ataccgcgag	catttccaac	1320
atgacatccc	aggaatattgt	ggagaaaacta	accaagcgac	tgaaaagaca	ccctgaggag	1380
acaggaggct	tccaggaggc	accgctggcc	tatgatgcca	tctgggcctt	ggcactggcc	1440
ctgaacaaga	catctggagg	aggcggccgt	tctggtgtgc	gcctggagga	cttcaactac	1500
aacaaccaga	ccattaccga	ccaaatctac	cgggcaatga	actcttcgtc	ctttgagggg	1560
gtctctggcc	atgtggtggt	tgatgccagc	ggctctcgga	tggcatggac	gcttatcgag	1620
cagcttcagg	gtggcagcta	caagaagatt	ggctactatg	acagcaccaa	ggatgatctt	1680
tcctgggtcca	aaacagataa	atggattgga	gggtccccc	cagctgacca	gacctgggtc	1740
atcaagacat	tccgcttcc	gtcacagaaa	ctctttatct	ccgtctcagt	tctctccagc	1800
ctgggcattg	tccatagctgt	tgtctgtctg	tcctttaaca	tctacaactc	acatgtccgt	1860
tatatccaga	actcacagcc	caacctgaac	aacctgactg	ctgtgggctg	ctcactggct	1920
ttagctgctg	tcttccccct	ggggctcgat	ggttaccaca	ttgggaggaa	ccagtttcc	1980
ttcgtctgcc	aggcccgct	ctggctcctg	ggcctgggct	ttagtctggg	ctacgggtcc	2040
atgttcacca	agatttggtg	ggccacacg	gtcttcacaa	agaaggaaga	aaagaaggag	2100
tggaggaaga	ctctggaacc	ctggaagctg	tatgccacag	tgggcctgct	ggtgggcatg	2160
gatgtcctca	ctctcgccat	ctggcagatc	gtggaccctc	tgcaccggac	cattgagaca	2220
tttgccaagg	aggaacctaa	ggaagatatt	gacgtctcta	ttctgcccc	gctggagcat	2280
tgcagctcca	ggaagatgaa	tacatggctt	ggcattttct	atgggttaca	ggggctgctg	2340
ctgctgctgg	gaatcttcc	tgcattatgag	accaagagtg	tgtccactga	gaagatcaat	2400
gatcaccggg	ctgtgggcat	ggctatctac	aatgtggcag	tctgtgcct	catcactgct	2460
cctgtcacca	tgattctgtc	cagccagcag	gatgcagcct	ttgcctttgc	ctctcttgcc	2520
atagttttct	cctcctatat	cactcttggt	gtgctctttg	tgcccaagat	gcgcaggctg	2580
atcacccgag	gggaatggca	gtcggaggcg	caggacacca	tgaagacagg	gtcatcgacc	2640
aacaacaacg	aggagagaaa	gtcccggctg	ttggagaagg	agaaccgtga	actggaaaag	2700
atcattgctg	agaaagagga	gcgtgtctct	gaactgcgcc	atcagctcca	gtctcggcag	2760
cagctccgct	cccggcgcca	cccaccgaca	cccccagaac	cctctggggg	cctgcccagg	2820
ggacccccctg	agccccccga	ccggcttagc	tgtgatggga	gtcagtgca	tttgctttat	2880
aagtgagggt	agggtgaggg	aggacaggcc	agtaggggga	gggaaaggga	gaggggaagg	2940
gcaggggact	caggaagcag	ggggctccca	tcccagctg	ggaagaacat	gctatccaat	3000
ctcatctctt	gtaatacat	gtccccctgt	gagttctggg	ctgatttggg	tctctcatac	3060
ctctgggaaa	cagacctttt	tctctcttac	tgcttcatgt	aattttgtat	cacctcttca	3120
caatttagtt	cgtacctggc	ttgaagctgc	tactgtctca	cacgtgcct	cctcagcagc	3180
ctcactgcat	ctttctcttc	ccatgcaaca	ccctcttcta	gttaccacgg	caaccctgc	3240
agctcctctg	cctttgtgct	ctgttcctgt	ccagcagggg	tctcccaaca	agtgtctttt	3300
ccaccccaaa	ggggcctctc	cttttctcca	ctgtcataat	ctctttccat	cttacttgcc	3360
cttctatact	ttctcacatg	tggctcccc	tgaattttgc	ttcctttggg	gagctcattc	3420
ntttcgccaa	ggntcacatg	ctcccttgcc	tctggctccg	tgca		3464

<210> 20  
 <211> 2887  
 <212> DNA  
 <213> Human

<400> 20

atggggccccg	gggccccctt	tgcccggtg	gggtggccac	tgcgccttct	ggttgtgatg	60
gcggcagggg	tggctccggt	gtgggcctcc	cactcccccc	atctcccgcg	gcctcactcg	120
cgggtcccc	cgcacccctc	ctcagaacgg	cgcgcagtgt	acatcggggc	actgtttccc	180
atgagcgggg	gctggccagg	gggcccaggc	tgccagcccc	cgggtggagat	ggcgtggag	240
gacgtgaata	gccgcaggga	catcctgccc	gactatgagc	tcaagctcat	ccaccacgac	300
agcaagtgtg	atccaggcca	agccaccaag	tacctatatg	agctgctcta	caacgaccct	360

atcaagatca	tccttatgcc	tggtgcagc	tctgtctcca	cgctggtggc	tgaggctgct	420
aggatgtgga	acctcattgt	gctttccat	ggctccagct	caccagccct	gtcaaaccgg	480
cagcgtttcc	ccactttctt	ccgaacgcac	ccatcagcca	cactccacaa	ccctaccggc	540
gtgaaactct	ttgaaaagt	gggctggaag	aagattgcta	ccatccagca	gaccactgag	600
gtcttcactt	cgactctgga	cgacctggag	gaacgagtga	aggaggctgg	aattgagatt	660
actttccgcc	agagtttctt	ctcagatcca	gctgtgccc	tcaaaaacct	gaagcgccag	720
gatgcccga	tcacgtgagg	acttttctat	gagactgaag	cccggaaagt	tttttgtgag	780
gtgtacaagg	agcgtctctt	tggaagaag	tacgtctggt	tcctcattgg	gtggtatgct	840
gacaattggt	tcaagatcta	cgaccttct	atcaactgca	cagtggatga	gatgactgag	900
gcgggtggag	gccacatcac	aactgagatt	gtcatgctga	atcctgccaa	taccgcgagc	960
atttccaaca	tgacatccca	ggaatttgtg	gagaaactaa	ccaagcgact	gaaaagacac	1020
cctgaggaga	caggaggctt	ccaggaggca	ccgctggcct	atgatgccat	ctgggccttg	1080
gcactggccc	tgaacaagac	atctggagga	ggcgggcgtt	ctgggtgtgc	cctggaggac	1140
ttcaactaca	acaaccagac	cattaccgac	caaactctacc	gggcaatgaa	ctcttcgtcc	1200
tttgagggtg	tctctggcca	tgtggtgttt	gatgccagcg	gctctcggat	ggcatggacg	1260
cttatcgagc	agcttcaggg	tggcagctac	aagaagattg	gctactatga	cagcaccaag	1320
gatgatcttt	cctggtccaa	aacagataaa	tggattggag	gggtcccccc	agctgaccag	1380
accctggtca	tcaagacatt	ccgcttcctg	tcacagaaac	tctttatctc	cgtctcagtt	1440
ctctccagcc	tgggcattgt	cctagctgtt	gtctgtctgt	cctttaacat	ctacaactca	1500
catgtccgtt	atatccagaa	ctcacagccc	aacctgaaca	acctgactgc	tgtgggctgc	1560
tcactggctt	tagctgctgt	cttccccctg	gggctcgatg	gttaccacat	tgggaggaac	1620
cagtttccct	tcgtctgcca	ggcccgctc	tggctcctgg	gcctgggctt	tagtctgggc	1680
tacggttcca	tgttcaccaa	gatttgggtg	gtccacacgg	tcttcacaaa	gaaggaagaa	1740
aagaaggagt	ggaggaagac	tctggaaccc	tggaaagctg	atgccacagt	gggcctgctg	1800
gtgggcatgg	atgtcctcac	tctcgccatc	tggcagatcg	tggacctct	gcaccggacc	1860
attgagacat	ttgccaaagga	ggaacctaa	gaagatattg	acgtctctat	tctgccccag	1920
ctggagcatt	gcagctccag	gaagatgaat	acatggcttg	gcattttcta	tggttacaag	1980
gggctgctgc	tgctgctggg	aatcttcctt	gcttatgaga	ccaagagtgt	gtccactgag	2040
aagatcaatg	atcaccgggc	tgtgggcatg	gctatctaca	atgtggcagt	cctgtgcctc	2100
atcactgctc	ctgtcaccat	gattctgtcc	agccagcagg	atgcagcctt	tgcctttgcc	2160
tctcttgcca	tagttttctc	ctcctatata	actcttgttg	tgctctttgt	gccccagatg	2220
cgcaggctga	tcacccgagg	ggaatggcag	tcggaggcgc	aggacaccat	gaagacaggg	2280
tcacgcacca	acaacaacga	ggaggagaag	tcccggtgt	tggagaagga	gaaccgtgaa	2340
ctggaaaaga	tcattgtctga	gaaagaggag	cgtgtctctg	aactgcgcca	tcaactccag	2400
tctcggcagc	agctccgctc	ccggcgccac	ccaccgacac	ccccagaacc	ctctgggggc	2460
ctgcccagg	gacccctga	gcccccgac	cggcttagct	gtgatgggag	tcgagtgcac	2520
ttgctttata	agtgagggtg	gggtgaggga	ggacaggcca	gtagggggag	ggaaagggag	2580
aggggaagg	caggggactc	aggaagcagg	gggtcccat	ccccagctgg	gaagaacatg	2640
ctatccaatc	tcactctctg	taaatacatg	tccccctgtg	agttctgggc	tgatttgggt	2700
ctctcatacc	tctgggaaac	agacctttt	ctctcttact	gcttcatgta	attttgtatc	2760
acctcttcac	aatttagttc	gtacctggct	tgaagctgct	cactgctcac	acgtgcctc	2820
ctcagcagcc	tcactgcac	tttctcttcc	catgcaacac	cctcttctag	ttaccacggc	2880
aacccct						2887

&lt;210&gt; 21

&lt;211&gt; 3144

&lt;212&gt; DNA

&lt;213&gt; Human

&lt;400&gt; 21

atggcttccc	cgcggagctc	cgggcagccc	gggcccngc	cgccgcgccc	accgcccgcc	60
gcgcgcctgc	tactgctact	gctgctgccg	ctgctgctgc	ctctggcgcc	cggggcctgg	120
ggctgggcgc	ggggcgcccc	ccggccgcgc	cccagcagcc	cgccgctctc	catcatgggc	180
ctcatgccgc	tcaccaagga	ggtggccaag	ggcagcatcg	ggcgcggtgt	gctccccgcc	240

gtggaactgg	ccatcgagca	gatccgcaac	gagtcactcc	tgcgccccta	cttcctcgac	300
ctgcggtctt	atgacacgga	gtgcgacaac	gcaaaagggt	tgaaagcctt	ctacgatgca	360
ataaaatacg	ggccgaacca	cttgatggtg	tttggaggcg	tctgtccatc	cgtcacatcc	420
atcattgcag	agtcctccca	aggctggaat	ctggtgcagc	tttcttttgc	tgcaaccacg	480
cctgttctag	ccgataagaa	aaaataccct	tatttctttc	ggaccgtccc	atcagacaat	540
gcggtgaatc	cagccattct	gaagttgctc	aagcactacc	agtggaagcg	cgtgggcacg	600
ctgacgcaag	acgttcagag	gttctctgag	gtgcggaatg	acctgactgg	agttctgtat	660
ggcgaggaca	ttgagatttc	agacaccgag	agcttctcca	acgatccctg	taccagtgtc	720
aaaaagctga	aggggaatga	tgtgcggatc	atccttggcc	agtttgacca	gaatatggca	780
gcaaaagtgt	tctgttgtgc	atacgaggag	aacatgtatg	gtagtaaata	tcagtggatc	840
attccgggct	ggtacgagcc	ttcttggtgg	gagcagggtg	acacggaagc	caactcatcc	900
cgctgcctcc	ggaagaatct	gcttgcctgc	atggagggtc	acattggcgt	ggatttcgag	960
cccctgagct	ccaagcagat	caagaccatc	tcaggaaaga	ctccacagca	gtatgagaga	1020
gagtacaaca	acaagcggtc	aggcgtgggg	ccagcaagt	tccacgggta	cgcttacgat	1080
ggcatctggg	tcatcgccaa	gacactgcag	agggccatgg	agacactgca	tgccagcagc	1140
cggcaccagc	ggatccagga	cttcaactac	acggaccaca	cgctgggcag	gatcatcctc	1200
aatgccatga	acgagacca	cttcttcggg	gtcacgggtc	aagttgtatt	ccggaatggg	1260
gagagaatgg	ggaccattaa	atttactcaa	tttcaagaca	gcagggaggt	gaaggtggga	1320
gagtacaacg	ctgtggccga	cacactggag	atcatcaatg	acaccatcag	gttccaagga	1380
tccgaaccac	caaaagacaa	gaccatcatc	ctggagcagc	tgcggaagat	ctccctacct	1440
ctctacagca	tctctctctg	cctcaccatc	ctcgggatga	tcatggccag	tgcttttctc	1500
ttcttcaaca	tcaagaaccg	gaatcagaag	ctcataaaga	tgtcagatcc	atacatgaac	1560
aaccttatca	tcttggagg	gatgctctcc	tatgcttcca	tatttctctt	tggccttgat	1620
ggatcctttg	tctctgaaaa	gacctttgaa	acactttgca	ccgtcaggac	ctggattctc	1680
accgtgggct	acacgaccgc	ttttggggcc	atgtttgcaa	agacctggag	agtccacgcc	1740
atcttcaaaa	atgtgaaaat	gaagaagaag	atcatcaagg	accagaaact	gcttgtgatc	1800
gtggggggcg	tgtgtctgat	cgacctgtgt	atcctgatct	gctggcaggc	tgtggacccc	1860
ctgcgaagga	cagtggagaa	gtacagcatg	gagccggacc	cagcaggacg	ggatatctcc	1920
atccgccctc	tctggagca	ctgtgagaac	acccatatga	ccatctggct	tggcatcgtc	1980
tatgcctaca	agggacttct	catgttgctt	ggttgcttct	tagcttggga	gacccgcaac	2040
gtcagcatcc	ccgactcaa	cgacagcaag	tacatcggga	tgagtgtcta	caacgtgggg	2100
atcatgtgca	tcatcggggc	cgtgtctctc	ttcctgacct	gggaccagcc	caatgtgcag	2160
ttctgcatcg	tggtcttggt	catcatcttc	tgcagacca	tcacctctg	cctggtattc	2220
gtgccgaagc	tcatcaccct	gagaacaaac	ccagatgcag	caacgcagaa	caggcgattc	2280
cagttcactc	agaatcagaa	gaaagaagat	tctaaaacgt	ccacctcggt	caccagtgtg	2340
aaccaagcca	gcacatcccc	cctggagggg	ctacagtcag	aaaaccatcg	cctgcgaatg	2400
aagatcacag	agctggataa	agacttgga	gaggtcacca	tgcagctgca	ggacacacca	2460
gaaaagacca	cctacattaa	acagaaccac	taccaagagc	tcaatgacat	cctcaacctg	2520
ggaaacttca	ctgagagcac	agatggagga	aaggccattt	taaaaaatca	cctcgatcaa	2580
aatccccagc	tacagtggaa	cacaacagag	ccctctcgaa	catgcaaaga	tcctatagaa	2640
gatataaact	ctccagaaca	catccagcgt	cggctgtccc	tccagctccc	catcctccac	2700
cacgcctacc	tcccatccat	cggaggcgtg	gacgccagct	gtgtcagccc	ctgcgtcagc	2760
cccaccgcca	gcccccgcca	cagacatgtg	ccaccctcct	tccgagtcac	ggtctcgggc	2820
ctgtaagggg	gggaggcctg	ggccccgggc	ctcccccggt	acagaaccac	actgggcaga	2880
ggggctctgt	gcgaaaacac	tgtcggctct	ggctgcggag	aagctgggca	ccatggctgg	2940
cctctcagga	ccactcggat	ggcactcagg	tggacaggac	ggggcagggg	gagacttggc	3000
acctgacctc	gagccttatt	tgtgaagtcc	ttatttcttc	acaaagaaga	ggaacggaaa	3060
tgggacgtct	tccttaacat	ctgcaaacaa	ggaggcgctg	ggatatcaaa	cttgcaaaaa	3120
aaaaaaaaaa	aaaaaaaaaa	aaaa				3144



<210> 22  
 <211> 2880  
 <212> DNA  
 <213> Rat

<400> 22

atgctgctgc	tgctgctggt	gcctctcttc	ctccgcccc	tgggcgctgg	cggggcgcag	60
accccccaacg	ccacctcgga	aggttgccag	attatacatc	cgccctggga	aggtggcatc	120
aggtaccgtg	gcttgactcg	cgaccagggtg	aaggccatca	acttcctgcc	tgtggactat	180
gagatcgaat	atgtgtgccc	aggggagcgc	gaggtggtgg	ggcccaaggt	gcgcaaatgc	240
ctggccaacg	gctcctggac	ggatatggac	acaccagcc	gctgtgtccg	aatctgctcc	300
aagtcttatt	tgaccttgga	aaatgggaag	gttttctga	cggttgggga	cctcccagct	360
ctggatggag	cccgggtgga	gttccgatgt	gaccccgact	tccatctggt	gggcagctcc	420
cggagcgtct	gtagtcaggg	ccagtgagagc	acccccaaagc	ccactgccca	ggtgaatcga	480
acgccacact	cagaacggcg	tgcagtatac	atcggggcgc	tgtttcccat	gagcgggggc	540
tggccggggg	gccaggcctg	ccagcccgcg	gtggagatgg	cgctggagga	cgtaaacagc	600
cgcagagaca	tcctgccgga	ctacgagctc	aagcttatcc	accacgacag	caagtgtgac	660
ccagggcaag	ccaccaagta	cttgtagcga	ctactctaca	atgaccccat	caagatcatt	720
ctcatgcctg	gctgtagttc	tgtctccaca	cttgtagctg	aggctgccc	gatgtggaac	780
cttattgtgc	tctcatatgg	ctccagttca	ccagccttgt	caaaccgaca	gcggtttccc	840
acgttcttcc	ggacgcattc	atccgccaca	ctccacaatc	ccaccgggt	gaaactcttc	900
gaaaagtggg	gctggaagaa	gatcgctacc	atccaacaga	ccaccgaggt	cttcacctca	960
acgctggatg	acctggagga	gcgagtgaag	gaggtgggga	tcgagatcac	tttccgacag	1020
agtttcttct	cggatccagc	tgtgcctggt	aaaaacctga	agcgtcaaga	tgctcgaatc	1080
atcgtagggac	ttttctatga	gacggaagcc	cggaaagttt	tttgtgaggt	ctataaggaa	1140
aggctctttg	ggaagaagta	cgctctggtc	ctcatcggtt	ggtagtctga	caactggttc	1200
aagacctatg	acccgtcaat	caattgtaca	gtggaagaaa	tgaccgaggg	ggtggagggc	1260
cacatcacca	cggagattgt	catgctgaac	cctgccaaaca	cccgaagcat	ttccaacatg	1320
acgtcacagg	aatttgtgga	gaaactaacc	aagcggctga	aaagacaccc	cgaggagact	1380
ggaggcttcc	aggaggcacc	actggcctat	gatgctatct	gggccttggc	tttggccttg	1440
aacaagacgt	ctggaggagg	tggctcgttc	ggcgtgcgcc	tggaggactt	taactacaac	1500
aaccagacca	ttacagacca	gatctaccgg	gccatgaact	cctcctcctt	tgagggcggt	1560
tctggccatg	tggctcttga	tgccagcggc	tcccggatgg	catggacact	tatcgagcag	1620
ctacagggcg	gcagctacaa	gaagatcggc	tactacgaca	gcaccaagga	tgatctttcc	1680
tgggtccaaaa	cggacaagtg	gattggaggg	tctccccag	ctgaccagac	cttggctcatc	1740
aagacattcc	gttttctgtc	tcagaaactc	tttatctccg	tctcagttct	ctccagcctg	1800
ggcattgttc	ttgctgttgt	ctgtctgtcc	tttaacatct	acaactccca	cgttcgttat	1860
atccagaact	cccagcccaa	cctgaacaat	ctgactgctg	tgggctgctc	actggcactg	1920
gctgctgtct	tccctctcgg	gctggatggg	taccacatag	ggagaagcca	gttcccgttt	1980
gtctgccagg	cccgcctttg	gctcttgggc	ttgggcttta	gtctgggcta	tggctctatg	2040
ttcaccaaga	tctggtgggt	ccacacagtc	ttcacgaaga	aggaggagaa	gaaggagtgg	2100
aggaagaccc	tagagccctg	gaaactctat	gccactgtgg	gcctgctggt	gggcattggat	2160
gtcctgactc	ttgccatctg	gcagattgtg	gaccccttgc	accgaaccat	tgagactttt	2220
gccaaggagg	aaccaaagga	agacatcgat	gtctccattc	tgccccagtt	ggagcactgc	2280
agctccaaga	agatgaatac	gtggcttggc	attttctatg	gttacaaggg	gctgctgctg	2340
ctgctgggaa	tctttcttgc	ttacgaaacc	aagagcgtgt	ccactgaaaa	gatcaatgac	2400
cacagggccg	tgggcatggc	tatctacaat	gtcgcggtcc	tgtgtctcat	cactgctcct	2460
gtgaccatga	tcttttccag	tcagcaggac	gcagcctttg	cctttgcctc	tctggccatc	2520
gtgttctctt	cctacatcac	tctggttgtg	ctctttgtgc	ccaagatgcg	caggctgatc	2580
acccgagggg	aatggcagtc	tgaaacgcag	gacaccatga	aaacaggatc	atccaccaac	2640
aacaacgagg	aagagaagtgc	ccgactgttg	gagaaggaaa	accgagaact	ggaaaagatc	2700
atcgctgaga	aagaggagcg	cgtctctgaa	ctgcgccatc	agctccagtc	tcggcagcaa	2760
ctccgctcac	ggcgccaccc	cccaacaccc	ccagatccct	ctgggggcct	tcccagggga	2820
ccctctgagc	cccctgaccg	gcttagctgt	gatgggagtc	gagtacattt	gctttacaag	2880

<210> 23  
 <211> 2532  
 <212> DNA  
 <213> Rat

<400> 23

atggggcccg	ggggaccctg	tacccagtg	gggtggccgc	tgccctcttct	gctggtgatg	60
gcggctgggg	tggtcccggt	gtgggcctct	cactccctc	atctcccgcg	gcctcaccgc	120
agggcccccc	cgcacccctc	ctcagaacgg	cgtgcagtat	acatcggggc	gctgtttccc	180
atgagcgggg	gctggccggg	gggccaggcc	tgccagcccg	cggtggagat	ggcgctggag	240
gacgttaaca	gccgcagaga	catcctgccg	gactacgagc	tcaagcttat	ccaccacgac	300
agcaagtgtg	accaggggca	agccaccaag	tacttgtagc	aactactcta	caatgacccc	360
atcaagatca	ttctcatgcc	tggtcttagt	tctgtctcca	cacttgtagc	tgaggctgcc	420
cggatgtgga	accttattgt	gctctcatat	ggctccagtt	caccagcctt	gtcaaaccga	480
cagcggtttc	ccacgtttctt	ccggacgcat	ccatccgcca	cactccacaa	ttccaccggg	540
gtgaaactct	tcgaaaagtg	gggctggaag	aagatcgcta	ccatccaaca	gaccaccgag	600
gtcttcacct	caacgctgga	tgacctggag	gagcgagtga	aagaggctgg	gatcgagatc	660
actttccgac	agagttttctt	ctcggatcca	gctgtgcctg	ttaaaaacct	gaagcgtcaa	720
gatgctcgaa	tcatcgtggg	acttttctat	gagacgggaag	cccggaaggt	tttttgtgag	780
gtctataagg	aaaggctctt	tggaagaag	tacgtctggg	tcctcatcgg	gtggtatgct	840
gacaactggg	tcaagacctt	tgacctgcca	atcaattgta	cagtgggaaga	aatgaccgag	900
gcggtgaggg	gccacatcac	cacggagatt	gtcatgctga	acctgcca	caccggaagc	960
atttccaaca	tgacgtcaca	ggaatttggt	gagaaactaa	ccaagcggct	gaaaagacac	1020
cccgaggaga	ctggaggctt	ccaggaggca	ccactggcct	atgatgctat	ctgggccttg	1080
gctttggcct	tgaacaagac	gtctggagga	ggtggtcggt	ccggcgtagc	cctggaggag	1140
tttaactaca	acaaccagac	cattacagac	cagatctacc	gggccatgaa	ctcctcctcc	1200
tttgaggggc	tttctggcca	tgtggtcttt	gatgccagcg	gctcccggat	ggcatggaca	1260
cttatcgagc	agctacaggg	cggcagctac	aagaagatcg	gctactacga	cagcaccaag	1320
gatgatcttt	cctggtccaa	aacggacaag	tggattggag	ggtctcccc	agctgaccag	1380
accttgggtc	tcaagacatt	ccgtttcctg	tctcagaaac	tctttatctc	cgtctcagtt	1440
ctctccagcc	tgggcattgt	tcttgctgtt	gtctgtctgt	cctttaacat	ctacaactcc	1500
cacgttcggt	atatccagaa	ctcccagccc	aacctgaaca	atctgactgc	tgtgggctgc	1560
tcactggcac	tggtctgtgt	cttccctctc	gggctggatg	gttaccacat	agggagaagc	1620
cagttcccg	ttgtctgcca	ggcccgccct	tggtctttgg	gcttgggctt	tagtctgggg	1680
tatggctcta	tggtcaccaa	gatctggtgg	gtccacacag	tcttcacgaa	gaaggaggag	1740
aagaaggagt	ggaggaagac	cctagagccc	tggaaactct	atgccactgt	gggcctgctg	1800
gtgggcatgg	atgtcctgac	tcttgccatc	tggcagattg	tggaacctt	gcaccgaacc	1860
attgagactt	ttgccaaagg	ggaaccaaag	gaagacatcg	atgtctccat	tctgccccag	1920
ttggagcact	gcagctccaa	gaagatgaat	acgtggcttg	gcattttcta	tggttacaag	1980
gggtgctgc	tgctgctggg	aatctttctt	gcttacgaaa	ccaagagcgt	gtccactgaa	2040
aagatcaatg	accacagggc	cgtgggcatg	gctatctaca	atgtcgcggt	cctgtgtctc	2100
atcactgctc	ctgtgacctt	gatcctttcc	agtcagcagg	acgcagcctt	tgcttttgcc	2160
tctctggcca	tcgtgttctc	ttctacatc	actctggttg	tgctctttgt	gcccagatg	2220
cgcaggctga	tcacccgagg	ggaatggcag	tctgaaacgc	aggacacat	gaaaacagga	2280
tcacccacca	acaacaacga	ggaagagaag	tcccgaactg	tggagaagga	aaaccgagaa	2340
ctggaaaaga	tcacgctgga	gaaagaggag	cgcgtctctg	aactgcgcca	tcagctccag	2400
tctcggcagc	aactccgctc	acggcgccac	cccccaacac	ccccagatcc	ctctgggggg	2460
cttcccaggg	gacctcttga	gccccctgac	cggcttagct	gtgatgggag	tcgagtacat	2520
ttgctttaca	ag					2532

<210> 24  
 <211> 960  
 <212> PRT  
 <213> Rat

<400> 24

Met	Leu	Leu	Leu	Leu	Leu	Val	Pro	Leu	Phe	Leu	Arg	Pro	Leu	Gly	Ala	
1				5					10					15		
Gly	Gly	Ala	Gln	Thr	Pro	Asn	Ala	Thr	Ser	Glu	Gly	Cys	Gln	Ile	Ile	
			20					25					30			
His	Pro	Pro	Trp	Glu	Gly	Gly	Ile	Arg	Tyr	Arg	Gly	Leu	Thr	Arg	Asp	
		35					40					45				
Gln	Val	Lys	Ala	Ile	Asn	Phe	Leu	Pro	Val	Asp	Tyr	Glu	Ile	Glu	Tyr	
	50					55					60					
Val	Cys	Arg	Gly	Glu	Arg	Glu	Val	Val	Gly	Pro	Lys	Val	Arg	Lys	Cys	
65					70					75					80	
Leu	Ala	Asn	Gly	Ser	Trp	Thr	Asp	Met	Asp	Thr	Pro	Ser	Arg	Cys	Val	
				85					90					95		
Arg	Ile	Cys	Ser	Lys	Ser	Tyr	Leu	Thr	Leu	Glu	Asn	Gly	Lys	Val	Phe	
			100					105					110			
Leu	Thr	Gly	Gly	Asp	Leu	Pro	Ala	Leu	Asp	Gly	Ala	Arg	Val	Glu	Phe	
		115					120					125				
Arg	Cys	Asp	Pro	Asp	Phe	His	Leu	Val	Gly	Ser	Ser	Arg	Ser	Val	Cys	
	130					135					140					
Ser	Gln	Gly	Gln	Trp	Ser	Thr	Pro	Lys	Pro	His	Cys	Gln	Val	Asn	Arg	
145				150						155					160	
Thr	Pro	His	Ser	Glu	Arg	Arg	Ala	Val	Tyr	Ile	Gly	Ala	Leu	Phe	Pro	
			165						170					175		
Met	Ser	Gly	Gly	Trp	Pro	Gly	Gly	Gln	Ala	Cys	Gln	Pro	Ala	Val	Glu	
		180					185						190			
Met	Ala	Leu	Glu	Asp	Val	Asn	Ser	Arg	Arg	Asp	Ile	Leu	Pro	Asp	Tyr	
		195					200					205				
Glu	Leu	Lys	Leu	Ile	His	His	Asp	Ser	Lys	Cys	Asp	Pro	Gly	Gln	Ala	
	210					215					220					
Thr	Lys	Tyr	Leu	Tyr	Glu	Leu	Leu	Tyr	Asn	Asp	Pro	Ile	Lys	Ile	Ile	
225				230						235					240	
Leu	Met	Pro	Gly	Cys	Ser	Ser	Val	Ser	Thr	Leu	Val	Ala	Glu	Ala	Ala	
			245						250					255		
Arg	Met	Trp	Asn	Leu	Ile	Val	Leu	Ser	Tyr	Gly	Ser	Ser	Ser	Pro	Ala	
			260					265						270		

Leu Ser Asn Arg Gln Arg Phe Pro Thr Phe Phe Arg Thr His Pro Ser  
 275 280 285  
 Ala Thr Leu His Asn Pro Thr Arg Val Lys Leu Phe Glu Lys Trp Gly  
 290 295 300  
 Trp Lys Lys Ile Ala Thr Ile Gln Gln Thr Thr Glu Val Phe Thr Ser  
 305 310 315 320  
 Thr Leu Asp Asp Leu Glu Glu Arg Val Lys Glu Ala Gly Ile Glu Ile  
 325 330 335  
 Thr Phe Arg Gln Ser Phe Phe Ser Asp Pro Ala Val Pro Val Lys Asn  
 340 345 350  
 Leu Lys Arg Gln Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu Thr  
 355 360 365  
 Glu Ala Arg Lys Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe Gly  
 370 375 380  
 Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp Phe  
 385 390 395 400  
 Lys Thr Tyr Asp Pro Ser Ile Asn Cys Thr Val Glu Glu Met Thr Glu  
 405 410 415  
 Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro Ala  
 420 425 430  
 Asn Thr Arg Ser Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu Lys  
 435 440 445  
 Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe Gln  
 450 455 460  
 Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala Leu  
 465 470 475 480  
 Asn Lys Thr Ser Gly Gly Gly Gly Arg Ser Gly Val Arg Leu Glu Asp  
 485 490 495  
 Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala Met  
 500 505 510  
 Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp Ala  
 515 520 525  
 Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly Gly  
 530 535 540  
 Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu Ser  
 545 550 555 560  
 Trp Ser Lys Thr Asp Lys Trp Ile Gly Gly Ser Pro Pro Ala Asp Gln  
 565 570 575

Ile Leu Val Ile Lys Thr Phe Arg Phe Leu Ser Gln Lys Leu Phe Ile  
 580 585 590  
 Ser Val Ser Val Leu Ser Ser Leu Gly Ile Val Leu Ala Val Val Cys  
 595 600 605  
 Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile Gln Asn Ser  
 610 615 620  
 Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala Leu  
 625 630 635 640  
 Ala Ala Val Phe Pro Leu Gly Leu Asp Gly Tyr His Ile Gly Arg Ser  
 645 650 655  
 Gln Phe Pro Phe Val Cys Gln Ala Arg Leu Trp Leu Leu Gly Leu Gly  
 660 665 670  
 Phe Ser Leu Gly Tyr Gly Ser Met Phe Thr Lys Ile Trp Trp Val His  
 675 680 685  
 Thr Val Phe Thr Lys Lys Glu Glu Lys Lys Glu Trp Arg Lys Thr Leu  
 690 695 700  
 Glu Pro Trp Lys Leu Tyr Ala Thr Val Gly Leu Leu Val Gly Met Asp  
 705 710 715 720  
 Val Leu Thr Leu Ala Ile Trp Gln Ile Val Asp Pro Leu His Arg Thr  
 725 730 735  
 Ile Glu Thr Phe Ala Lys Glu Glu Pro Lys Glu Asp Ile Asp Val Ser  
 740 745 750  
 Ile Leu Pro Gln Leu Glu His Cys Ser Ser Lys Lys Met Asn Thr Trp  
 755 760 765  
 Leu Gly Ile Phe Tyr Gly Tyr Lys Gly Leu Leu Leu Leu Gly Ile  
 770 775 780  
 Phe Leu Ala Tyr Glu Thr Lys Ser Val Ser Thr Glu Lys Ile Asn Asp  
 785 790 795 800  
 His Arg Ala Val Gly Met Ala Ile Tyr Asn Val Ala Val Leu Cys Leu  
 805 810 815  
 Ile Thr Ala Pro Val Thr Met Ile Leu Ser Ser Gln Gln Asp Ala Ala  
 820 825 830  
 Phe Ala Phe Ala Ser Leu Ala Ile Val Phe Ser Ser Tyr Ile Thr Leu  
 835 840 845  
 Val Val Leu Phe Val Pro Lys Met Arg Arg Leu Ile Thr Arg Gly Glu  
 850 855 860  
 Trp Gln Ser Glu Thr Gln Asp Thr Met Lys Thr Gly Ser Ser Thr Asn

[illegible]

Met 1	Gly	Pro	Gly	Gly 5	Pro	Cys	Thr	Pro	Val 10	Gly	Trp	Pro	Leu	Pro 15	Leu
Leu	Leu	Val	Met 20	Ala	Ala	Gly	Val	Ala 25	Pro	Val	Trp	Ala	Ser 30	His	Ser
Pro	His	Leu 35	Pro	Arg	Pro	His	Pro 40	Arg	Val	Pro	Pro	His 45	Pro	Ser	Ser
Glu	Arg	Arg	Ala	Val	Tyr	Ile 55	Gly	Ala	Leu	Phe	Pro 60	Met	Ser	Gly	Gly
Trp 65	Pro	Gly	Gly	Gln 70	Ala	Cys	Gln	Pro	Ala	Val 75	Glu	Met	Ala	Leu	Glu 80
Asp	Val	Asn	Ser	Arg 85	Arg	Asp	Ile	Leu 90	Pro	Asp	Tyr	Glu	Leu	Lys 95	Leu
Ile	His	His	Asp 100	Ser	Lys	Cys	Asp	Pro 105	Gly	Gln	Ala	Thr	Lys 110	Tyr	Leu
Tyr	Glu	Leu 115	Leu	Tyr	Asn	Asp	Pro 120	Ile	Lys	Ile	Ile	Leu 125	Met	Pro	Gly
Cys	Ser 130	Ser	Val	Ser	Thr	Leu 135	Val	Ala	Glu	Ala	Ala	Arg	Met	Trp	Asn
Leu 145	Ile	Val	Leu	Ser 150	Tyr	Gly	Ser	Ser	Ser	Pro 155	Ala	Leu	Ser	Asn	Arg 160
Gln	Arg	Phe	Pro	Thr	Phe	Phe	Arg	Thr	His	Pro	Ser	Ala	Thr	Leu	His

[illegible]

Lys Thr Phe Arg Phe Leu Ser Gln Lys Leu Phe Ile Ser Val Ser Val  
 465 470 475 480  
 Leu Ser Ser Leu Gly Ile Val Leu Ala Val Val Cys Leu Ser Phe Asn  
 485 490 495  
 Ile Tyr Asn Ser His Val Arg Tyr Ile Gln Asn Ser Gln Pro Asn Leu  
 500 505 510  
 Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala Leu Ala Ala Val Phe  
 515 520 525  
 Pro Leu Gly Leu Asp Gly Tyr His Ile Gly Arg Ser Gln Phe Pro Phe  
 530 535 540  
 Val Cys Gln Ala Arg Leu Trp Leu Leu Gly Leu Gly Phe Ser Leu Gly  
 545 550 555 560  
 Tyr Gly Ser Met Phe Thr Lys Ile Trp Trp Val His Thr Val Phe Thr  
 565 570 575  
 Lys Lys Glu Glu Lys Lys Glu Trp Arg Lys Thr Leu Glu Pro Trp Lys  
 580 585 590  
 Leu Tyr Ala Thr Val Gly Leu Leu Val Gly Met Asp Val Leu Thr Leu  
 595 600 605  
 Ala Ile Trp Gln Ile Val Asp Pro Leu His Arg Thr Ile Glu Thr Phe  
 610 615 620  
 Ala Lys Glu Glu Pro Lys Glu Asp Ile Asp Val Ser Ile Leu Pro Gln  
 625 630 635 640  
 Leu Glu His Cys Ser Ser Lys Lys Met Asn Thr Trp Leu Gly Ile Phe  
 645 650 655  
 Tyr Gly Tyr Lys Gly Leu Leu Leu Leu Leu Gly Ile Phe Leu Ala Tyr  
 660 665 670  
 Glu Thr Lys Ser Val Ser Thr Glu Lys Ile Asn Asp His Arg Ala Val  
 675 680 685  
 Gly Met Ala Ile Tyr Asn Val Ala Val Leu Cys Leu Ile Thr Ala Pro  
 690 695 700  
 Val Thr Met Ile Leu Ser Ser Gln Gln Asp Ala Ala Phe Ala Phe Ala  
 705 710 715 720  
 Ser Leu Ala Ile Val Phe Ser Ser Tyr Ile Thr Leu Val Val Leu Phe  
 725 730 735  
 Val Pro Lys Met Arg Arg Leu Ile Thr Arg Gly Glu Trp Gln Ser Glu  
 740 745 750  
 Thr Gln Asp Thr Met Lys Thr Gly Ser Ser Thr Asn Asn Asn Glu Glu



755	760	765
Glu Lys Ser Arg Leu Leu Glu Lys Glu Asn Arg Glu Leu Glu Lys Ile		
770	775	780
Ile Ala Glu Lys Glu Glu Arg Val Ser Glu Leu Arg His Gln Leu Gln		
785	790	795 800
Ser Arg Gln Gln Leu Arg Ser Arg Arg His Pro Pro Thr Pro Pro Asp		
805	810	815
Pro Ser Gly Gly Leu Pro Arg Gly Pro Ser Glu Pro Pro Asp Arg Leu		
820	825	830
Ser Cys Asp Gly Ser Arg Val His Leu Leu Tyr Lys		
835	840	

<210> 26  
 <211> 2616  
 <212> DNA  
 <213> Human

<400> 26

atgggatcgc	tgcttgctgt	cctggcactg	ctgccgctgt	ggggtgctgt	ggctgagggc	60
ccagccaaga	aggtgctgac	cctggaggga	gacttggtgc	tgggtgggct	gttcccagtg	120
caccagaagg	gcggcccagc	agaggactgt	ggctctgtca	atgagcaccg	tggcatccag	180
cgcttgagg	ccatgctttt	tgcactggac	cgcatcaacc	gtgaccgca	cctgctgcct	240
ggcgtgcgcc	tgggtgcaca	catcctcgac	agttgtcca	aggacacaca	tgcgtggag	300
caggcactgg	actttgtgcg	tgcctcactc	agcgtggtg	ctgatggatc	acgccacatc	360
tgccccgacg	gctcttatgc	gacccatggg	gatgtccca	ctgccatcac	tgggtgttatt	420
ggcggttcct	acagtgatgt	ctccatccag	gtggccaacc	tcttgaggct	atttcagatc	480
ccacagatta	gctacgcctc	taccagtgcc	aagctgagtg	acaagtcccg	ctatgactac	540
tttgcccga	cagtgcctcc	tgacttcttc	caagccaagg	ccatggctga	gattctccgc	600
ttcttcaact	ggacctatgt	gtccactgag	gcctctgagg	gcgactatgg	cgagacaggc	660
attgaggcct	ttgagctaga	ggctcgtgcc	cgcaacatct	gtgtggccac	ctcggagaaa	720
gtgggcccgtg	ccatgagccg	cgcgcccttt	gaggggtgtg	tgcgagccct	gctgcagaag	780
cccagtgcgc	gcgtggctgt	cctgttcacc	cgctctgagg	atgcccggga	gctgcttgct	840
gccagccagc	gcctcaatgc	cagcttcacc	tgggtggcca	gtgatggttg	gggggccctg	900
gagagtgtgg	tggcaggcag	tgagggggct	gctgaggggtg	ctatcaccat	cgagctggcc	960
tcctaccca	tcagtgaact	tgcctcctac	ttccagagcc	tggacccttg	gaacaacagc	1020
cggaacccct	ggttccgtga	attctgggag	cagagggttc	gctgcagctt	ccggcagcga	1080
gactgcgcag	cccactctct	ccgggctgtg	ccctttgaac	aggagtccaa	gatcatgttt	1140
gtggtcaatg	cagtgtacgc	catggcccat	gcgctccaca	acatgcaccg	tgccctctgc	1200
cccaacacca	cccggctctg	tgacgcgatg	cggccagtta	acgggcgcgc	cctctacaag	1260
gactttgtgc	tcaacgtcaa	gtttgatgcc	ccctttcgcc	cagctgacac	ccacaatgag	1320
gtccgctttg	accgcttttg	tgatggtatt	ggcgcgtaca	acatcttcac	ctatctgcgt	1380
gcaggcagtg	ggcgttatcg	ctaccagaag	gtgggctact	gggcagaagg	cttgactctg	1440
gacaccagcc	tcatcccatg	ggcctcacgc	tcagccggcc	ccctggccgc	ctctcgctgc	1500
agtgagccct	gcctccagaa	tgagggtgaag	agtgtgcagc	cgggcgaagt	ctgctgctgg	1560
ctctgcattc	cgtgccagcc	ctatgagtac	cgattggacg	aattcacttg	cgctgattgt	1620
ggcctgggct	actggcccaa	tgcagccctg	actggctgct	tcgaactgcc	ccaggagtac	1680
atccgctggg	gcgatgcctg	ggctgtggga	cctgtcacca	tcgcctgcct	cgggtgccctg	1740
gccaccctgt	ttgtgctggg	tgtctttgtg	cggcacaatg	ccacaccagt	gggtcaaggcc	1800
tcaggctcggg	agctctgcta	catcctgctg	gggtggtgtct	tcctctgcta	ctgcatgacc	1860
ttcatcttca	ttgccaaagg	atccacggca	gtgtgtacct	tacggcgtct	tgggttgggc	1920

```

actgccttct ctgtctgcta ctcagccctg ctcaccaaga ccaaccgcat tgcacgcattc 1980
ttcgggtgggg cccgggaggg tgcccagcgg ccacgcttca tcagtcctgc ctcacaggtg 2040
gccatctgcc tggcacttat ctcgggccag ctgctcatcg tggtcgcctg gctgggtggg 2100
gaggcaccgg gcacaggcaa ggagacagcc cccgaacggc gggaggtggt gacactgcgc 2160
tgcaaccacc gcgatgcaag tatgttgggc tcgctggcct acaatgtgct cctcatcgcg 2220
ctctgcacgc tttatgcctt caatactcgc aagtgcctcg aaaacttcaa cgaggccaag 2280
ttcattggct tcaccatgta caccacctgc atcatctggc tggcattgtt gcccatcttc 2340
tatgtcacct ccagtgacta ccgggtacag accaccacca tgtgcgtgtc agtcagcctc 2400
agcggctccg tgggtgcttg ctgcctcttt gcgcccagc tgcacatcat cctcttccag 2460
ccgcagaaga acgtgggttag ccaccgggca cccaccagcc gctttggcag tgctgctgcc 2520
agggccagct ccagccttgg ccaaggtctt ggctcccagt ttgtcccccac tgtttgcaat 2580
ggcgtgagg tgggtggactc gacaacgtca tcgctt 2616

```

```

<210> 27
<211> 824
<212> PRT
<213> Human

```

```

<400> 27

```

```

Met Gly Ser Leu Leu Ala Leu Leu Ala Leu Leu Pro Leu Trp Gly Ala
 1             5             10             15
Val Ala Glu Gly Pro Ala Lys Lys Val Leu Thr Leu Glu Gly Asp Leu
          20             25             30
Val Leu Gly Gly Leu Phe Pro Val His Gln Lys Gly Gly Pro Ala Glu
          35             40             45
Asp Cys Gly Pro Val Asn Glu His Arg Gly Ile Gln Arg Leu Glu Ala
 50             55             60
Met Leu Phe Ala Leu Asp Arg Ile Asn Arg Asp Pro His Leu Leu Pro
 65             70             75             80
Gly Val Arg Leu Gly Ala His Ile Leu Asp Ser Cys Ser Lys Asp Thr
          85             90             95
His Ala Leu Glu Gln Ala Leu Asp Phe Val Arg Ala Ser Leu Ser Arg
          100            105            110
Gly Ala Asp Gly Ser Arg His Ile Cys Pro Asp Gly Ser Tyr Ala Thr
          115            120            125
His Gly Asp Ala Pro Thr Ala Ile Thr Gly Val Ile Gly Gly Ser Tyr
          130            135            140
Ser Asp Val Ser Ile Gln Val Ala Asn Leu Leu Arg Leu Phe Gln Ile
 145            150            155            160
Pro Gln Ile Ser Tyr Ala Ser Thr Ser Ala Lys Leu Ser Asp Lys Ser
          165            170            175
Arg Tyr Asp Tyr Phe Ala Arg Thr Val Pro Pro Asp Phe Phe Gln Ala

```

180	185	190
Lys Ala Met Ala Glu Ile Leu Arg Phe Phe Asn Trp Thr Tyr Val Ser		
195	200	205
Thr Glu Ala Ser Glu Gly Asp Tyr Gly Glu Thr Gly Ile Glu Ala Phe		
210	215	220
Glu Leu Glu Ala Arg Ala Arg Asn Ile Cys Val Ala Thr Ser Glu Lys		
225	230	235
Val Gly Arg Ala Met Ser Arg Ala Ala Phe Glu Gly Val Val Arg Ala		
245	250	255
Leu Leu Gln Lys Pro Ser Ala Arg Val Ala Val Leu Phe Thr Arg Ser		
260	265	270
Glu Asp Ala Arg Glu Leu Leu Ala Ala Ser Gln Arg Leu Asn Ala Ser		
275	280	285
Phe Thr Trp Val Ala Ser Asp Gly Trp Gly Ala Leu Glu Ser Val Val		
290	295	300
Ala Gly Ser Glu Gly Ala Ala Glu Gly Ala Ile Thr Ile Glu Leu Ala		
305	310	315
Ser Tyr Pro Ile Ser Asp Phe Ala Ser Tyr Phe Gln Ser Leu Asp Pro		
325	330	335
Trp Asn Asn Ser Arg Asn Pro Trp Phe Arg Glu Phe Trp Glu Gln Arg		
340	345	350
Phe Arg Cys Ser Phe Arg Gln Arg Asp Cys Ala Ala His Ser Leu Arg		
355	360	365
Ala Val Pro Phe Glu Gln Glu Ser Lys Ile Met Phe Val Val Asn Ala		
370	375	380
Val Tyr Ala Met Ala His Ala Leu His Asn Met His Arg Ala Leu Cys		
385	390	395
Pro Asn Thr Thr Arg Leu Cys Asp Ala Met Arg Pro Val Asn Gly Arg		
405	410	415
Arg Leu Tyr Lys Asp Phe Val Leu Asn Val Lys Phe Asp Ala Pro Phe		
420	425	430
Arg Pro Ala Asp Thr His Asn Glu Val Arg Phe Asp Arg Phe Gly Asp		
435	440	445
Gly Ile Gly Arg Tyr Asn Ile Phe Thr Tyr Leu Arg Ala Gly Ser Gly		
450	455	460
Arg Tyr Arg Tyr Gln Lys Val Gly Tyr Trp Ala Glu Gly Leu Thr Leu		
465	470	475
Asp Thr Ser Leu Ile Pro Trp Ala Ser Pro Ser Ala Gly Pro Leu Ala		

				485				490				495			
Ala	Ser	Arg	Cys	Ser	Glu	Pro	Cys	Leu	Gln	Asn	Glu	Val	Lys	Ser	Val
			500				505						510		
Gln	Pro	Gly	Glu	Val	Cys	Cys	Trp	Leu	Cys	Ile	Pro	Cys	Gln	Pro	Tyr
			515				520				525				
Glu	Tyr	Arg	Leu	Asp	Glu	Phe	Thr	Cys	Ala	Asp	Cys	Gly	Leu	Gly	Tyr
			530				535				540				
Trp	Pro	Asn	Ala	Ser	Leu	Thr	Gly	Cys	Phe	Glu	Leu	Pro	Gln	Glu	Tyr
545				550						555			560		
Ile	Arg	Trp	Gly	Asp	Ala	Trp	Ala	Val	Gly	Pro	Val	Thr	Ile	Ala	Cys
			565						570			575			
Leu	Gly	Ala	Leu	Ala	Thr	Leu	Phe	Val	Leu	Gly	Val	Phe	Val	Arg	His
			580						585			590			
Asn	Ala	Thr	Pro	Val	Val	Lys	Ala	Ser	Gly	Arg	Glu	Leu	Cys	Tyr	Ile
			595			600						605			
Leu	Leu	Gly	Gly	Val	Phe	Leu	Cys	Tyr	Cys	Met	Thr	Phe	Ile	Phe	Ile
			610			615			620						
Ala	Lys	Pro	Ser	Thr	Ala	Val	Cys	Thr	Leu	Arg	Arg	Leu	Gly	Leu	Gly
625				630						635			640		
Thr	Ala	Phe	Ser	Val	Cys	Tyr	Ser	Ala	Leu	Leu	Thr	Lys	Thr	Asn	Arg
			645						650			655			
Ile	Ala	Arg	Ile	Phe	Gly	Gly	Ala	Arg	Glu	Gly	Ala	Gln	Arg	Pro	Arg
			660			665						670			
Phe	Ile	Ser	Pro	Ala	Ser	Gln	Val	Ala	Ile	Cys	Leu	Ala	Leu	Ile	Ser
			675			680						685			
Gly	Gln	Leu	Leu	Ile	Val	Val	Ala	Trp	Leu	Val	Val	Glu	Ala	Pro	Gly
			690			695			700						
Thr	Gly	Lys	Glu	Thr	Ala	Pro	Glu	Arg	Arg	Glu	Val	Val	Thr	Leu	Arg
705				710						715			720		
Cys	Asn	His	Arg	Asp	Ala	Ser	Met	Leu	Gly	Ser	Leu	Ala	Tyr	Asn	Val
			725						730			735			
Leu	Leu	Ile	Ala	Leu	Cys	Thr	Leu	Tyr	Ala	Phe	Asn	Thr	Arg	Lys	Cys
			740			745						750			
Pro	Glu	Asn	Phe	Asn	Glu	Ala	Lys	Phe	Ile	Gly	Phe	Thr	Met	Tyr	Thr
			755			760						765			
Thr	Cys	Ile	Ile	Trp	Leu	Ala	Leu	Leu	Pro	Ile	Phe	Tyr	Val	Thr	Ser
			770			775			780						

Ser Asp Tyr Arg Val Gln Thr Thr Thr Met Cys Val Ser Val Ser Leu  
785 790 795 800

Ser Gly Ser Val Val Leu Gly Cys Leu Phe Ala Pro Lys Leu His Ile  
805 810 815

Ile Leu Phe Gln Pro Gln Lys Asn  
820

<210> 28  
<211> 1077  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Chimeric Gqi5

<400> 28

atgactctgg	agtccatcat	ggcgtgctgc	ctgagcgagg	aggccaagga	agcccggcgg	60
atcaacgacg	agatcgagcg	gcagctccgc	agggacaagc	gggacgcccg	ccgggagctc	120
aagctgctgc	tgctcgggac	aggagagagt	ggcaagagta	cgtttatcaa	gcagatgaga	180
atcatccatg	ggtcaggata	ctctgatgaa	gataaaaagg	gcttcaccaa	gctgggtgat	240
cagaacatct	tcacggccat	gcaggccatg	atcagagcca	tggacacact	caagatccca	300
tacaagtatg	agcacaataa	ggctcatgca	caattagttc	gagaagttga	tgtggagaag	360
gtgtctgctt	ttgagaatcc	atatgtagat	gcaataaaga	gtttatggaa	tgatcctgga	420
atccaggaat	gctatgatag	acgacgagaa	tatcaattat	ctgactctac	caaatactat	480
cttaatgact	tggaccgctg	agctgaccct	gcctacctgc	ctacgcaaca	agatgtgctt	540
agagtctcag	tccccaccac	agggatcatc	gaatacccct	ttgacttaca	aagtgtcatt	600
ttcagaatgg	tcgatgtagg	gggccaaagg	tcagagagaa	gaaaatggat	acactgcttt	660
gaaaatgtca	cctctatcat	gtttctagta	gcgcttagtg	aatatgatca	agttctcgtg	720
gagtcagaca	atgagaaccg	aatggaggaa	agcaaggctc	tctttagaac	aattatcaca	780
tacccttggt	tccagaactc	ctcggttatt	ctgttcttaa	acaagaaaga	tcttctagag	840
gagaaaatca	tgtattccca	tctagtcgac	tacttcccag	aatatgatgg	acccagagag	900
gatgccacgg	cagcccagag	attcattctg	aagatgttcg	tggacctgaa	cccagacagt	960
gacaaaatta	tctactccca	cttcacgtgc	gccacagaca	ccgagaatat	ccgctttgtc	1020
tttgctgccg	tcaaggacac	catectccag	ttgaacctga	aggactgcgg	tctgttc	1077

<210> 29  
<211> 359  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Chimeric Gqi5

<400> 29

Met Thr Leu Glu Ser Ile Met Ala Cys Cys Leu Ser Glu Glu Ala Lys  
1 5 10 15

Glu Ala Arg Arg Ile Asn Asp Glu Ile Glu Arg Gln Leu Arg Arg Asp  
20 25 30

Lys Arg Asp Ala Arg Arg Glu Leu Lys Leu Leu Leu Gly Thr Gly

35					40					45					
Glu	Ser	Gly	Lys	Ser	Thr	Phe	Ile	Lys	Gln	Met	Arg	Ile	Ile	His	Gly
50						55					60				
Ser	Gly	Tyr	Ser	Asp	Glu	Asp	Lys	Arg	Gly	Phe	Thr	Lys	Leu	Val	Tyr
65					70					75					80
Gln	Asn	Ile	Phe	Thr	Ala	Met	Gln	Ala	Met	Ile	Arg	Ala	Met	Asp	Thr
				85					90					95	
Leu	Lys	Ile	Pro	Tyr	Lys	Tyr	Glu	His	Asn	Lys	Ala	His	Ala	Gln	Leu
			100					105					110		
Val	Arg	Glu	Val	Asp	Val	Glu	Lys	Val	Ser	Ala	Phe	Glu	Asn	Pro	Tyr
		115					120						125		
Val	Asp	Ala	Ile	Lys	Ser	Leu	Trp	Asn	Asp	Pro	Gly	Ile	Gln	Glu	Cys
	130						135				140				
Tyr	Asp	Arg	Arg	Arg	Glu	Tyr	Gln	Leu	Ser	Asp	Ser	Thr	Lys	Tyr	Tyr
145					150					155					160
Leu	Asn	Asp	Leu	Asp	Arg	Val	Ala	Asp	Pro	Ala	Tyr	Leu	Pro	Thr	Gln
				165					170					175	
Gln	Asp	Val	Leu	Arg	Val	Arg	Val	Pro	Thr	Thr	Gly	Ile	Ile	Glu	Tyr
			180					185						190	
Pro	Phe	Asp	Leu	Gln	Ser	Val	Ile	Phe	Arg	Met	Val	Asp	Val	Gly	Gly
		195					200					205			
Gln	Arg	Ser	Glu	Arg	Arg	Lys	Trp	Ile	His	Cys	Phe	Glu	Asn	Val	Thr
	210					215					220				
Ser	Ile	Met	Phe	Leu	Val	Ala	Leu	Ser	Glu	Tyr	Asp	Gln	Val	Leu	Val
225					230					235					240
Glu	Ser	Asp	Asn	Glu	Asn	Arg	Met	Glu	Glu	Ser	Lys	Ala	Leu	Phe	Arg
				245					250					255	
Thr	Ile	Ile	Thr	Tyr	Pro	Trp	Phe	Gln	Asn	Ser	Ser	Val	Ile	Leu	Phe
			260					265						270	
Leu	Asn	Lys	Lys	Asp	Leu	Leu	Glu	Glu	Lys	Ile	Met	Tyr	Ser	His	Leu
		275					280					285			
Val	Asp	Tyr	Phe	Pro	Glu	Tyr	Asp	Gly	Pro	Gln	Arg	Asp	Ala	Gln	Ala
	290					295					300				
Ala	Arg	Glu	Phe	Ile	Leu	Lys	Met	Phe	Val	Asp	Leu	Asn	Pro	Asp	Ser
305					310					315					320
Asp	Lys	Ile	Ile	Tyr	Ser	His	Phe	Thr	Cys	Ala	Thr	Asp	Thr	Glu	Asn
				325					330					335	
Ile	Arg	Phe	Val	Phe	Ala	Ala	Val	Lys	Asp	Thr	Ile	Leu	Gln	Leu	Asn

340

345

350

Leu Lys Asp Cys Gly Leu Phe  
355

&lt;210&gt; 30

&lt;211&gt; 2751

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; Chimeric hCAR/hmGluR2

&lt;400&gt; 30

atggcatttt	atagctgctg	ctgggtcctc	ttggcactca	cctggcacac	ctctgcctac	60
gggccagacc	agcgagccca	aaagaagggg	gacattatcc	ttggggggct	ctttcctatt	120
cattttggag	tagcagctaa	agatcaagat	ctcaaatcaa	ggccggagtc	tgtggaatgt	180
atcagggtata	atttccgtgg	gtttcgtctg	ttacaggcta	tgatatttgc	catagaggag	240
ataaacagca	gcccagccct	tcttccaac	ttgacgctgg	gatacaggat	atgtgacact	300
tgcaacaccg	tttctaaggc	cttgggaagcc	accctgagtt	ttgttgctca	aaacaaaatt	360
gattctttga	accttgatga	gttctgcaac	tgctcagagc	acattccctc	tacgattgct	420
gtggtgggag	caactggctc	aggcgtctcc	acggcagtg	caaactctgt	ggggctcttc	480
tacattcccc	aggctcagta	tgctcctccc	agcagactcc	tcagcaacaa	gaatcaattc	540
aagtctttcc	tccgaaccat	ccccaatgat	gagcaccagg	ccactgccat	ggcagacatc	600
atcgagtatt	tccgctggaa	ctgggtgggc	acaattgcag	ctgatgacga	ctatgggagg	660
ccggggattg	agaaattccg	agaggaagct	gaggaagagg	atatctgcat	cgacttcagt	720
gaactcatct	cccagtaact	tgatgaggaa	gagatccagc	atgtggtaga	ggtgattcaa	780
aattccacgg	ccaaagtcac	cgtggttttc	tccagtggcc	cagatcttga	gccccctatc	840
aaggagattg	tccggcgcaa	tatcacgggc	aagatctggc	tggccagcga	ggcctggggc	900
agctcctccc	tgatcgccat	gcctcagtac	ttccacgtgg	ttggcggcac	cattggattc	960
gctctgaagg	ctgggcagat	cccaggcttc	cgggaattcc	tgaagaaggt	ccatcccagg	1020
aagtctgtcc	acaatggttt	tgccaaggag	ttttgggaag	aaacatttaa	ctgccacctc	1080
caagaagggtg	caaaaggacc	tttacctgtg	gacacctttc	tgagagggtca	cgaagaaagt	1140
ggcgacagggt	ttagcaacag	ctcgacagcc	ttccgacccc	tctgtacagg	ggatgagaac	1200
atcagcagtg	tcgagacccc	ttacatagat	tacacgcatt	tacggatatc	ctacaatgtg	1260
tacttagcag	tctactccat	tgcccacgcc	ttgcaagata	tatatacctg	cttacctggg	1320
agagggctct	tcaccaatgg	ctcctgtgca	gacatcaaga	aagttgaggc	gtggcagggtc	1380
ctgaagcacc	tacggcatct	aaactttaca	aacaatatgg	gggagcagggt	gacctttgat	1440
gagtgtgggtg	acctgggtggg	gaactatttc	atcatcaact	ggcacctctc	cccagaggat	1500
ggctccatcg	tgtttaagga	agtcgggtat	tacaacgtct	atgccaagaa	gggagaaaga	1560
ctcttcatca	acgaggagaa	aatcctgtgg	agtgggttct	ccagggaggt	gcccttctcc	1620
aactgcagcc	gagactgcct	ggcagggacc	aggaaaggga	tcattgaggg	ggagcccacc	1680
tgctgctttg	agtgtgtgga	gtgtcctgat	ggggagtata	gtgatgagac	agatgccagt	1740
gcctgtaaca	agtgcccgaa	tgacttctgg	tccaatgaga	accacacctc	ctgcttcgaa	1800
ctgccccagg	agtacatccg	ctggggcgat	gcctgggctg	tgggacctgt	caccatcgcc	1860
tgctcgggtg	ccctggccac	cctgtttgtg	ctgggtgtct	ttgtgcggca	caatgccaca	1920
ccagtgggtca	aggcctcagg	tcgggagctc	tgctacatcc	tgctgggtgg	tgtcttctct	1980
tgctactgca	tgaccttcat	cttcattgcc	aagccatcca	cggcagtggtg	taccttacgg	2040
cgtcttggtt	tgggcactgc	cttctctgtc	tgctactcag	ccctgctcac	caagaccaac	2100
cgcatcgac	gcattcttcg	tggggcccgg	gagggtgccc	agcggccacg	cttcatcagt	2160
cctgcctcac	agggtggccat	ctgcctggca	cttatctcgg	gccagctgct	catcgtgggtc	2220
gcctggctgg	tgggtggaggc	accgggcaca	ggcaaggaga	cagccccgga	acggcggggag	2280
gtggtgacac	tgcgctgcaa	ccaccgcgat	gcaagtatgt	tgggctcgct	ggcctacaat	2340
gtgctcctca	tcgcgctctg	cacgctttat	gccttcaata	ctcgcaagtg	ccccgaaaac	2400
ttcaacgagg	ccaagttcat	tggcttcacc	atgtacacca	cctgcatcat	ctggctggga	2460

```

ttgttgccca ttttctatgt cacctccagt gactaccggg tacagaccac caccatgtgc 2520
gtgtcagtcg gcttcagcgg ctccgtgggtg cttggctgcc tctttgcgcc caagctgcac 2580
atcatcctct tccagccgca gaagaacgtg gttagccacc gggcaccac cagccgcttt 2640

ggcagtgtctg ctgccagggc cagctccagc cttggccaag ggtctggctc ccagtttgtc 2700
cccactgttt gcaatggcgc tgaggtgggtg gactcgacaa cgtcatcgct t 2751

```

&lt;210&gt; 31

&lt;211&gt; 917

&lt;212&gt; PRT

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; Chimeric hCAR/hmGluR2

&lt;400&gt; 31

```

Met Ala Phe Tyr Ser Cys Cys Trp Val Leu Leu Ala Leu Thr Trp His
 1              5              10              15

Thr Ser Ala Tyr Gly Pro Asp Gln Arg Ala Gln Lys Lys Gly Asp Ile
          20              25              30

Ile Leu Gly Gly Leu Phe Pro Ile His Phe Gly Val Ala Ala Lys Asp
      35              40              45

Gln Asp Leu Lys Ser Arg Pro Glu Ser Val Glu Cys Ile Arg Tyr Asn
      50              55              60

Phe Arg Gly Phe Arg Trp Leu Gln Ala Met Ile Phe Ala Ile Glu Glu
      65              70              75              80

Ile Asn Ser Ser Pro Ala Leu Leu Pro Asn Leu Thr Leu Gly Tyr Arg
          85              90              95

Ile Phe Asp Thr Cys Asn Thr Val Ser Lys Ala Leu Glu Ala Thr Leu
      100              105              110

Ser Phe Val Ala Gln Asn Lys Ile Asp Ser Leu Asn Leu Asp Glu Phe
      115              120              125

Cys Asn Cys Ser Glu His Ile Pro Ser Thr Ile Ala Val Val Gly Ala
      130              135              140

Thr Gly Ser Gly Val Ser Thr Ala Val Ala Asn Leu Leu Gly Leu Phe
      145              150              155              160

Tyr Ile Pro Gln Val Ser Tyr Ala Ser Ser Ser Arg Leu Leu Ser Asn
          165              170              175

Lys Asn Gln Phe Lys Ser Phe Leu Arg Thr Ile Pro Asn Asp Glu His
          180              185              190

Gln Ala Thr Ala Met Ala Asp Ile Ile Glu Tyr Phe Arg Trp Asn Trp
      195              200              205

```



Val Gly Thr Ile Ala Ala Asp Asp Asp Tyr Gly Arg Pro Gly Ile Glu  
 210 215 220  
 Lys Phe Arg Glu Glu Ala Glu Glu Arg Asp Ile Cys Ile Asp Phe Ser  
 225 230 235 240  
 Glu Leu Ile Ser Gln Tyr Ser Asp Glu Glu Glu Ile Gln His Val Val  
 245 250 255  
 Glu Val Ile Gln Asn Ser Thr Ala Lys Val Ile Val Val Phe Ser Ser  
 260 265 270  
 Gly Pro Asp Leu Glu Pro Leu Ile Lys Glu Ile Val Arg Arg Asn Ile  
 275 280 285  
 Thr Gly Lys Ile Trp Leu Ala Ser Glu Ala Trp Ala Ser Ser Ser Leu  
 290 295 300  
 Ile Ala Met Pro Gln Tyr Phe His Val Val Gly Gly Thr Ile Gly Phe  
 305 310 315 320  
 Ala Leu Lys Ala Gly Gln Ile Pro Gly Phe Arg Glu Phe Leu Lys Lys  
 325 330 335  
 Val His Pro Arg Lys Ser Val His Asn Gly Phe Ala Lys Glu Phe Trp  
 340 345 350  
 Glu Glu Thr Phe Asn Cys His Leu Gln Glu Gly Ala Lys Gly Pro Leu  
 355 360 365  
 Pro Val Asp Thr Phe Leu Arg Gly His Glu Glu Ser Gly Asp Arg Phe  
 370 375 380  
 Ser Asn Ser Ser Thr Ala Phe Arg Pro Leu Cys Thr Gly Asp Glu Asn  
 385 390 395 400  
 Ile Ser Ser Val Glu Thr Pro Tyr Ile Asp Tyr Thr His Leu Arg Ile  
 405 410 415  
 Ser Tyr Asn Val Tyr Leu Ala Val Tyr Ser Ile Ala His Ala Leu Gln  
 420 425 430  
 Asp Ile Tyr Thr Cys Leu Pro Gly Arg Gly Leu Phe Thr Asn Gly Ser  
 435 440 445  
 Cys Ala Asp Ile Lys Lys Val Glu Ala Trp Gln Val Leu Lys His Leu  
 450 455 460  
 Arg His Leu Asn Phe Thr Asn Asn Met Gly Glu Gln Val Thr Phe Asp  
 465 470 475 480  
 Glu Cys Gly Asp Leu Val Gly Asn Tyr Ser Ile Ile Asn Trp His Leu  
 485 490 495

Ser Pro Glu Asp Gly Ser Ile Val Phe Lys Glu Val Gly Tyr Tyr Asn  
 500 505 510  
 Val Tyr Ala Lys Lys Gly Glu Arg Leu Phe Ile Asn Glu Glu Lys Ile  
 515 520 525  
 Leu Trp Ser Gly Phe Ser Arg Glu Val Pro Phe Ser Asn Cys Ser Arg  
 530 535 540  
 Asp Cys Leu Ala Gly Thr Arg Lys Gly Ile Ile Glu Gly Glu Pro Thr  
 545 550 555 560  
 Cys Cys Phe Glu Cys Val Glu Cys Pro Asp Gly Glu Tyr Ser Asp Glu  
 565 570 575  
 Thr Asp Ala Ser Ala Cys Asn Lys Cys Pro Asp Asp Phe Trp Ser Asn  
 580 585 590  
 Glu Asn His Thr Ser Cys Phe Glu Leu Pro Gln Glu Tyr Ile Arg Trp  
 595 600 605  
 Gly Asp Ala Trp Ala Val Gly Pro Val Thr Ile Ala Cys Leu Gly Ala  
 610 615 620  
 Leu Ala Thr Leu Phe Val Leu Gly Val Phe Val Arg His Asn Ala Thr  
 625 630 635 640  
 Pro Val Val Lys Ala Ser Gly Arg Glu Leu Cys Tyr Ile Leu Leu Gly  
 645 650 655  
 Gly Val Phe Leu Cys Tyr Cys Met Thr Phe Ile Phe Ile Ala Lys Pro  
 660 665 670  
 Ser Thr Ala Val Cys Thr Leu Arg Arg Leu Gly Leu Gly Thr Ala Phe  
 675 680 685  
 Ser Val Cys Tyr Ser Ala Leu Leu Thr Lys Thr Asn Arg Ile Ala Arg  
 690 695 700  
 Ile Phe Gly Gly Ala Arg Glu Gly Ala Gln Arg Pro Arg Phe Ile Ser  
 705 710 715 720  
 Pro Ala Ser Gln Val Ala Ile Cys Leu Ala Leu Ile Ser Gly Gln Leu  
 725 730 735  
 Leu Ile Val Val Ala Trp Leu Val Val Glu Ala Pro Gly Thr Gly Lys  
 740 745 750  
 Glu Thr Ala Pro Glu Arg Arg Glu Val Val Thr Leu Arg Cys Asn His  
 755 760 765  
 Arg Asp Ala Ser Met Leu Gly Ser Leu Ala Tyr Asn Val Leu Leu Ile  
 770 775 780  
 Ala Leu Cys Thr Leu Tyr Ala Phe Asn Thr Arg Lys Cys Pro Glu Asn  
 785 790 795 800

Phe Asn Glu Ala Lys Phe Ile Gly Phe Thr Met Tyr Thr Thr Cys Ile  
                     805                    810                    815  
 Ile Trp Leu Ala Leu Leu Pro Ile Phe Tyr Val Thr Ser Ser Asp Tyr  
                     820                    825                    830  
 Arg Val Gln Thr Thr Thr Met Cys Val Ser Val Ser Leu Ser Gly Ser  
                     835                    840                    845  
 Val Val Leu Gly Cys Leu Phe Ala Pro Lys Leu His Ile Ile Leu Phe  
                     850                    855                    860  
 Gln Pro Gln Lys Asn Val Val Ser His Arg Ala Pro Thr Ser Arg Phe  
                     865                    870                    875                    880  
 Gly Ser Ala Ala Ala Arg Ala Ser Ser Ser Leu Gly Gln Gly Ser Gly  
                     885                    890                    895  
 Ser Gln Phe Val Pro Thr Val Cys Asn Gly Arg Glu Val Val Asp Ser  
                     900                    905                    910  
 Thr Thr Ser Ser Leu  
                     915

&lt;210&gt; 32

&lt;211&gt; 3831

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; Chimeric phCaR/hmGluR2\*Gqi5

&lt;400&gt; 32

atggcatttt	atagctgctg	ctgggtcctc	ttggcactca	cctggcacac	ctctgcctac	60
gggccagacc	agcgagccca	aaagaagggg	gacattatcc	ttggggggct	ctttcctatt	120
cattttggag	tagcagctaa	agatcaagat	ctcaaataca	ggccggagtc	tgtggaatgt	180
atcaggtata	atttccgtgg	gtttcgctgg	ttacaggcta	tgatatttgc	catagaggag	240
ataaacagca	gcccagccct	tcttcccaac	ttgacgctgg	gatacaggat	atttgacact	300
tgcaacaccg	tttctaaggc	cttggaagcc	acctgagtt	ttgttgctca	aaacaaaatt	360
gattctttga	accttgatga	gttctgcaac	tgctcagagc	acattccctc	tacgattgct	420
gtgggtgggag	caactggctc	aggcgtctcc	acggcagtg	caaactctgt	ggggctcttc	480
tacattcccc	aggtcagtta	tgctctctcc	agcagactcc	tcagcaacaa	gaatcaattc	540
aagtctttcc	tccgaaccat	ccccaatgat	gagcaccagg	ccactgccat	ggcagacatc	600
atcgagtatt	tccgctggaa	ctgggtgggc	acaattgcag	ctgatgacga	ctatggggcg	660
ccggggattg	agaaattccg	agaggaagct	gaggaaaggg	atatctgcat	cgacttcagt	720
gaactcatct	cccagtactc	tgatgaggaa	gagatccagc	atgtggtaga	ggtgattcaa	780
aattccacgg	caaagtcatt	cgtgggtttc	tccagtggcc	cagatcttga	gcccctcatc	840
aaggagattg	tccggcgcaa	tatcacgggc	aagatctggc	tggccagcga	ggcctggggc	900
agctctctcc	tgatcgccat	gcctcagtac	ttccacgtgg	ttggcggcac	cattggattc	960
gctctgaagg	ctgggcagat	cccaggcttc	cggaattcc	tgaagaaggt	ccatcccagg	1020
aagtctgtcc	acaatggttt	tgccaaggag	ttttgggaag	aaacatttaa	ctgccacctc	1080
caagaaggtg	caaaaggacc	tttacctgtg	gacacctttc	tgagaggtca	cgaagaaagt	1140

ggcgacaggt	ttagcaacag	ctcgacagcc	ttccgacccc	tctgtacagg	ggatgagaac	1200
atcagcagtg	tcgagacccc	ttacatagat	tacacgcatt	tacggatatc	ctacaatgtg	1260
tacttagcag	tctactccat	tgcccacgcc	ttgcaagata	tatataacctg	cttacctggg	1320
agagggctct	tcaccaatgg	ctcctgtgca	gacatcaaga	aagttgaggc	gtggcaggtc	1380
ctgaagcacc	tacggcatct	aaactttaca	aacaatatgg	gggagcaggt	gacctttgat	1440
gagtgtggtg	acctggtggg	gaactattcc	atcatcaact	ggcacctctc	cccagaggat	1500
ggctccatcg	tgtttaagga	agtcggtat	tacaacgtct	atgccaagaa	gggagaaaga	1560
ctcttcatca	acgaggagaa	aatcctgtgg	agtgggttct	ccagggagggt	gcccttctcc	1620
aactgcagcc	gagactgcct	ggcagggacc	aggaaggga	tcattgaggg	ggagcccacc	1680
tgtctgtttg	agtgtgtgga	gtgtccgtat	ggggagtata	gtgatgagac	agatgccagt	1740
gcctgtaaca	agtgcccaga	tgacttctgg	tccaatgaga	accacacctc	ctgtctcgaa	1800
ctgccccagg	agtacatccg	ctggggcgat	gcctgggctg	tgggacctgt	caccatcgcc	1860
tgctcggtg	ccctggccac	cctgtttgtg	ctgggtgtct	ttgtgcgga	caatgccaca	1920
ccagtggtea	aggcctcagg	tcgggagctc	tgctacatcc	tgtctgggtg	tgtcttctc	1980
tgctactgca	tgaccttcat	cttcattgcc	aagccatcca	cggcagtggt	taccttacgg	2040
cgtcttggtt	tgggcactgc	cttctctgtc	tgctactcag	cctgtctcac	caagaccaac	2100
cgatttgca	gcattctcgg	tggggcccgg	gagggtgccc	agcgccaccg	cttcatcagt	2160
cctgcctcac	aggtggccat	ctgcctggca	cttatctcgg	gccagctgct	catcgtggtc	2220
gcctggctgg	tggtggaggc	accgggcaca	ggcaaggaga	cagccccgga	acggcgggag	2280
gtggtgacac	tgcgctgcaa	ccaccgcgat	gcaagtatgt	tgggctcgct	ggcctacaat	2340
gtgctctcca	tgcgcctctg	cacgctttat	gccttcaata	ctcgcaagtg	ccccgaaac	2400
ttcaacgagg	ccaagttcat	tggcttcacc	atgtacacca	cctgcacat	ctggctggca	2460
ttgttgccca	tcttctatgt	cacctccagt	gactaccggg	tacagaccac	caccatgtgc	2520
gtgtcagtea	gcctcagcgg	ctccgtgggtg	cttggctgcc	tctttgcgcc	caagctgcac	2580
atcatcctct	tccagccgca	gaagaacgtg	gttagccacc	gggcaccacc	cagccgcttt	2640
ggcagtgctg	ctgccagggc	cagctccagc	cttggccaag	ggtctggctc	ccagtttgtc	2700
ccactggtt	gcaatggccg	tgaggtgggtg	gactcgacaa	cgcatcgct	tatgactctg	2760
gagtcacatca	tggcgtgctg	cctgagcgag	gaggccaagg	aagcccggcg	gatcaacgac	2820
gagatcgagc	ggcagctccg	cagggacaag	cgggacgccc	gccgggagct	caagctgctg	2880
ctgctcggga	caggagagag	tggcaagagt	acgtttatca	agcagatgag	aatcatccat	2940
gggtcaggat	actctgatga	agataaaaag	ggcttcacca	agctgggtga	tcagaacatc	3000
ttcacggcca	tgcaggccat	gatacagacc	atggacacac	tcaagatccc	atacaagtat	3060
gagcacaata	aggctactgc	acaattagtt	ctgaaagtgt	atgtggagaa	ggtgtctgct	3120
tttgagaatc	catatgtaga	tgcaataaag	agtttatgga	atgatectgg	aatccaggaa	3180
tgctatgata	gacgacgaga	atatcaatta	tctgactcta	ccaaatacta	tcttaatgac	3240
ttggaccgcg	tagctgaccc	tgcttacctg	cctacgcaac	aagatgtgct	tagagttcga	3300
gtccccacca	cagggatcat	cgaatacccc	tttgacttac	aaagtgtcat	tttcagaatg	3360
gtcgatgtag	ggggccaaag	gtcagagaga	agaaaatgga	tacactgctt	tgaaaatgtc	3420
acctctatca	tgtttctagt	agcgcttagt	gaatatgata	aagtttctgt	ggagtcagac	3480
aatgagaacc	gaatggagga	aagcaaggct	ctcttttagaa	caattatcac	atacccttgg	3540
ttccagaaet	cctcggttat	tctgtttctta	aacaagaaag	atcttctaga	ggagaaaatc	3600
atgtattccc	atctagtcca	ctaactccca	gaatatgatg	gaccccagag	agatggccag	3660
gcagcccgcg	aattcattct	gaagatgttc	gtggacata	accagacag	tgacaaaatt	3720
atcatctccc	acttcacgtg	cgccacagac	accgagaata	tccgctttgt	ctttgtgtcc	3780
gtcaaggaca	ccatcctcca	gttgaacctg	aaggactgcg	gtctgttcta	a	3831

<210> 33  
 <211> 1276  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Chimeric phCaR/hmGluR2\*Gqi5  
  
 <400> 33

Met	Ala	Phe	Tyr	Ser	Cys	Cys	Trp	Val	Leu	Leu	Ala	Leu	Thr	Trp	His	1	5	10	15
Thr	Ser	Ala	Tyr	Gly	Pro	Asp	Gln	Arg	Ala	Gln	Lys	Lys	Gly	Asp	Ile	20	25	30	
Ile	Leu	Gly	Gly	Leu	Phe	Pro	Ile	His	Phe	Gly	Val	Ala	Ala	Lys	Asp	35	40	45	
Gln	Asp	Leu	Lys	Ser	Arg	Pro	Glu	Ser	Val	Glu	Cys	Ile	Arg	Tyr	Asn	50	55	60	
Phe	Arg	Gly	Phe	Arg	Trp	Leu	Gln	Ala	Met	Ile	Phe	Ala	Ile	Glu	Glu	65	70	75	80
Ile	Asn	Ser	Ser	Pro	Ala	Leu	Leu	Pro	Asn	Leu	Thr	Leu	Gly	Tyr	Arg	85	90	95	
Ile	Phe	Asp	Thr	Cys	Asn	Thr	Val	Ser	Lys	Ala	Leu	Glu	Ala	Thr	Leu	100	105	110	
Ser	Phe	Val	Ala	Gln	Asn	Lys	Ile	Asp	Ser	Leu	Asn	Leu	Asp	Glu	Phe	115	120	125	
Cys	Asn	Cys	Ser	Glu	His	Ile	Pro	Ser	Thr	Ile	Ala	Val	Val	Gly	Ala	130	135	140	
Thr	Gly	Ser	Gly	Val	Ser	Thr	Ala	Val	Ala	Asn	Leu	Leu	Gly	Leu	Phe	145	150	155	160
Tyr	Ile	Pro	Gln	Val	Ser	Tyr	Ala	Ser	Ser	Ser	Arg	Leu	Leu	Ser	Asn	165	170	175	
Lys	Asn	Gln	Phe	Lys	Ser	Phe	Leu	Arg	Thr	Ile	Pro	Asn	Asp	Glu	His	180	185	190	
Gln	Ala	Thr	Ala	Met	Ala	Asp	Ile	Ile	Glu	Tyr	Phe	Arg	Trp	Asn	Trp	195	200	205	
Val	Gly	Thr	Ile	Ala	Ala	Asp	Asp	Asp	Tyr	Gly	Arg	Pro	Gly	Ile	Glu	210	215	220	
Lys	Phe	Arg	Glu	Glu	Ala	Glu	Glu	Arg	Asp	Ile	Cys	Ile	Asp	Phe	Ser	225	230	235	240
Glu	Leu	Ile	Ser	Gln	Tyr	Ser	Asp	Glu	Glu	Glu	Ile	Gln	His	Val	Val	245	250	255	

Glu Val Ile Gln Asn Ser Thr Ala Lys Val Ile Val Val Phe Ser Ser  
 260 265 270  
 Gly Pro Asp Leu Glu Pro Leu Ile Lys Glu Ile Val Arg Arg Asn Ile  
 275 280 285  
 Thr Gly Lys Ile Trp Leu Ala Ser Glu Ala Trp Ala Ser Ser Ser Leu  
 290 295 300  
 Ile Ala Met Pro Gln Tyr Phe His Val Val Gly Gly Thr Ile Gly Phe  
 305 310 315 320  
 Ala Leu Lys Ala Gly Gln Ile Pro Gly Phe Arg Glu Phe Leu Lys Lys  
 325 330 335  
 Val His Pro Arg Lys Ser Val His Asn Gly Phe Ala Lys Glu Phe Trp  
 340 345 350  
 Glu Glu Thr Phe Asn Cys His Leu Gln Glu Gly Ala Lys Gly Pro Leu  
 355 360 365  
 Pro Val Asp Thr Phe Leu Arg Gly His Glu Glu Ser Gly Asp Arg Phe  
 370 375 380  
 Ser Asn Ser Ser Thr Ala Phe Arg Pro Leu Cys Thr Gly Asp Glu Asn  
 385 390 395 400  
 Ile Ser Ser Val Glu Thr Pro Tyr Ile Asp Tyr Thr His Leu Arg Ile  
 405 410 415  
 Ser Tyr Asn Val Tyr Leu Ala Val Tyr Ser Ile Ala His Ala Leu Gln  
 420 425 430  
 Asp Ile Tyr Thr Cys Leu Pro Gly Arg Gly Leu Phe Thr Asn Gly Ser  
 435 440 445  
 Cys Ala Asp Ile Lys Lys Val Glu Ala Trp Gln Val Leu Lys His Leu  
 450 455 460  
 Arg His Leu Asn Phe Thr Asn Asn Met Gly Glu Gln Val Thr Phe Asp  
 465 470 475 480  
 Glu Cys Gly Asp Leu Val Gly Asn Tyr Ser Ile Ile Asn Trp His Leu  
 485 490 495  
 Ser Pro Glu Asp Gly Ser Ile Val Phe Lys Glu Val Gly Tyr Tyr Asn  
 500 505 510  
 Val Tyr Ala Lys Lys Gly Glu Arg Leu Phe Ile Asn Glu Glu Lys Ile  
 515 520 525  
 Leu Trp Ser Gly Phe Ser Arg Glu Val Pro Phe Ser Asn Cys Ser Arg  
 530 535 540  
 Asp Cys Leu Ala Gly Thr Arg Lys Gly Ile Ile Glu Gly Glu Pro Thr

545		550		555		560
Cys Cys Phe Glu Cys Val Glu Cys Pro Asp Gly Glu Tyr Ser Asp Glu						
		565		570		575
Thr Asp Ala Ser Ala Cys Asn Lys Cys Pro Asp Asp Phe Trp Ser Asn						
		580		585		590
Glu Asn His Thr Ser Cys Phe Glu Leu Pro Gln Glu Tyr Ile Arg Trp						
		595		600		605
Gly Asp Ala Trp Ala Val Gly Pro Val Thr Ile Ala Cys Leu Gly Ala						
		610		615		620
Leu Ala Thr Leu Phe Val Leu Gly Val Phe Val Arg His Asn Ala Thr						
		625		630		635
Pro Val Val Lys Ala Ser Gly Arg Glu Leu Cys Tyr Ile Leu Leu Gly						
		645		650		655
Gly Val Phe Leu Cys Tyr Cys Met Thr Phe Ile Phe Ile Ala Lys Pro						
		660		665		670
Ser Thr Ala Val Cys Thr Leu Arg Arg Leu Gly Leu Gly Thr Ala Phe						
		675		680		685
Ser Val Cys Tyr Ser Ala Leu Leu Thr Lys Thr Asn Arg Ile Ala Arg						
		690		695		700
Ile Phe Gly Gly Ala Arg Glu Gly Ala Gln Arg Pro Arg Phe Ile Ser						
		705		710		715
Pro Ala Ser Gln Val Ala Ile Cys Leu Ala Leu Ile Ser Gly Gln Leu						
		725		730		735
Leu Ile Val Val Ala Trp Leu Val Val Glu Ala Pro Gly Thr Gly Lys						
		740		745		750
Glu Thr Ala Pro Glu Arg Arg Glu Val Val Thr Leu Arg Cys Asn His						
		755		760		765
Arg Asp Ala Ser Met Leu Gly Ser Leu Ala Tyr Asn Val Leu Leu Ile						
		770		775		780
Ala Leu Cys Thr Leu Tyr Ala Phe Asn Thr Arg Lys Cys Pro Glu Asn						
		785		790		795
Phe Asn Glu Ala Lys Phe Ile Gly Phe Thr Met Tyr Thr Thr Cys Ile						
		805		810		815
Ile Trp Leu Ala Leu Leu Pro Ile Phe Tyr Val Thr Ser Ser Asp Tyr						
		820		825		830
Arg Val Gln Thr Thr Thr Met Cys Val Ser Val Ser Leu Ser Gly Ser						
		835		840		845

Val Val Leu Gly Cys Leu Phe Ala Pro Lys Leu His Ile Ile Leu Phe  
 850 855 860  
 Gln Pro Gln Lys Asn Val Val Ser His Arg Ala Pro Thr Ser Arg Phe  
 865 870 875 880  
 Gly Ser Ala Ala Ala Arg Ala Ser Ser Ser Leu Gly Gln Gly Ser Gly  
 885 890 895  
 Ser Gln Phe Val Pro Thr Val Cys Asn Gly Arg Glu Val Val Asp Ser  
 900 905 910  
 Thr Thr Ser Ser Leu Met Thr Leu Glu Ser Ile Met Ala Cys Cys Leu  
 915 920 925  
 Ser Glu Glu Ala Lys Glu Ala Arg Arg Ile Asn Asp Glu Ile Glu Arg  
 930 935 940  
 Gln Leu Arg Arg Asp Lys Arg Asp Ala Arg Arg Glu Leu Lys Leu Leu  
 945 950 955 960  
 Leu Leu Gly Thr Gly Glu Ser Gly Lys Ser Thr Phe Ile Lys Gln Met  
 965 970 975  
 Arg Ile Ile His Gly Ser Gly Tyr Ser Asp Glu Asp Lys Arg Gly Phe  
 980 985 990  
 Thr Lys Leu Val Tyr Gln Asn Ile Phe Thr Ala Met Gln Ala Met Ile  
 995 1000 1005  
 Arg Ala Met Asp Thr Leu Lys Ile Pro Tyr Lys Tyr Glu His Asn Lys  
 1010 1015 1020  
 Ala His Ala Gln Leu Val Arg Glu Val Asp Val Glu Lys Val Ser Ala  
 1025 1030 1035 1040  
 Phe Glu Asn Pro Tyr Val Asp Ala Ile Lys Ser Leu Trp Asn Asp Pro  
 1045 1050 1055  
 Gly Ile Gln Glu Cys Tyr Asp Arg Arg Arg Glu Tyr Gln Leu Ser Asp  
 1060 1065 1070  
 Ser Thr Lys Tyr Tyr Leu Asn Asp Leu Asp Arg Val Ala Asp Pro Ala  
 1075 1080 1085  
 Tyr Leu Pro Thr Gln Gln Asp Val Leu Arg Val Arg Val Pro Thr Thr  
 1090 1095 1100  
 Gly Ile Ile Glu Tyr Pro Phe Asp Leu Gln Ser Val Ile Phe Arg Met  
 1105 1110 1115 1120  
 Val Asp Val Gly Gly Gln Arg Ser Glu Arg Arg Lys Trp Ile His Cys  
 1125 1130 1135  
 Phe Glu Asn Val Thr Ser Ile Met Phe Leu Val Ala Leu Ser Glu Tyr  
 1140 1145 1150



```

Asp Gln Val Leu Val Glu Ser Asp Asn Glu Asn Arg Met Glu Glu Ser
    1155                      1160                      1165

Lys Ala Leu Phe Arg Thr Ile Ile Thr Tyr Pro Trp Phe Gln Asn Ser
    1170                      1175                      1180

Ser Val Ile Leu Phe Leu Asn Lys Lys Asp Leu Leu Glu Glu Lys Ile
    1185                      1190                      1195                      1200

Met Tyr Ser His Leu Val Asp Tyr Phe Pro Glu Tyr Asp Gly Pro Gln
    1205                      1210                      1215

Arg Asp Ala Gln Ala Ala Arg Glu Phe Ile Leu Lys Met Phe Val Asp
    1220                      1225                      1230

Leu Asn Pro Asp Ser Asp Lys Ile Ile Tyr Ser His Phe Thr Cys Ala
    1235                      1240                      1245

Thr Asp Thr Glu Asn Ile Arg Phe Val Phe Ala Ala Val Lys Asp Thr
    1250                      1255                      1260

Ile Leu Gln Leu Asn Leu Lys Asp Cys Gly Leu Phe
    1265                      1270                      1275

```

```

<210> 34
<211> 3105
<212> DNA
<213> Artificial Sequence

<220>
<223> Chimeric hmGluR2/hCaR

<400> 34

```

```

atgggatcgc tgcttgcgct cccggcactg ctgctgctgt ggggtgctgt ggctgagggc      60
ccagccaaga aggtgctgac cctggagggg gacttggtgc tgggtgggct gttcccagtg      120
caccagaagg gcggcccagc agaggactgt ggtcctgtca atgagcaccg tggcatccag      180
cgcttgagg ccattgctttt tgcactggac cgcatacaac gtgaccgcga cctgctgcct      240
ggcgtgcgcc tgggtgcaca catcctcgac agttgctcca aggacacaca tgcgctggag      300
caggcactgg actttgtgcg tgccctcact agcctggtg ctgatggctc acgccacatc      360
tgccccgacg gctcttatgc gacccatggt gatgctccca ctgccatcac tgggtgttatt      420
ggcggttcct acagtgatgt ctccatccag gtggccaacc tcttgaggct atttcagatc      480
ccacagatta gctacgcctc taccagtgcc aagctgagtg acaagtcccg ctatgactac      540
tttggccgca cagtgcctcc tgacttcttc caagccaagg ccatggctga gattctccgc      600
ttcttcaact ggacctatgt gtccactgtg gcgtctgagg gcgactatgg cgagacaggc      660
attgaggcct ttgagctaga ggctcgtgcc cgcaacatct gtgtggccac ctcgagaaaa      720
gtggggccgtg ccatgagccg cgcggccttt gaggggtgtg tgcgagccct gctgcagaag      780
cccagtggcc gcgtggctgt cctgttcacc cgttctgagg atgcccgga gctgcttgct      840
gccagccagc gcctcaatgc cagcttcacc tgggtggcca gtgatggtt gggggccctg      900
gagagtgtgg tggcaggcag tgagggggct gctgaggggt ctatcaccat cgagctggcc      960
tcctaccca tcagtgactt tgccctctac ttccagagcc tggacccttg gaacaacagc     1020
cggaaccctt ggttccgtga attctgggag cagaggttcc gctgcagctt ccggcagcga     1080
gactgcgcag cccactctct ccgggctgtg ccctttgagc aggagtccaa gatcatgttt     1140

```

```

gtggtcaatg cagtgtacgc catggcccat gcgctccaca acatgcaccg tgcctctctgc 1200
cccaacacca cccggctctg tgacgcgatg cggccagtta acgggcgcgc cctctacaag 1260
gacttttgtg tcaacgtcaa gtttgatgcc ccctttcgcc cagctgacac ccacaatgag 1320
gtccgctttg accgcttttg tgatggtatt ggccgctaca acatcttcac ctatctcgct 1380
gcaggcagtg ggcgctatcg ctaccagaag gtgggctact gggcagaagg cttgactctg 1440
gacaccagcc tcatcccatg ggccctaccc tcagccggcc ccctgcccgc ctctcgctgc 1500
agtgagccct gctccagaa tgaggtgaag agtgtgcagc cgggcgaagt ctgctgctgg 1560
ctctgcattc cgtgccagcc ctatgagtac cgattggacg aattcacttg cgctgattgt 1620
ggcctgggct actggcccaa tgccagcctg actggctgct tcgaactgcc ccaggagtac 1680
atccgctggg gcgatgcctg ggctgtggga cctgtcacca tcgcctgcct cggtgccctg 1740
gccaccctct ttgtgctggg tgtctttgtg cggcacaatg ccacaccagt ggtcaaggcc 1800
tcaggtcggg agctctgcta catcctgctg ggtggtgtct tcctctgcta ctgcatgacc 1860
ttcatcttca ttgccaagcc atccacggca gtgtgtacct tacggcgtct tggtttgggc 1920
actgccttct ctgtctgcta ctacgcctg ctcaccaaga ccaaccgcat tgcacgcac 1980
ttcggtgagg cccgggaggg tgcccagcgg ccacgcttca tcagtcctgc ctcacagggtg 2040
gccatctgcc tggcacttat ctggggccag ctgctcatcg tggtcgcctg gctggtggtg 2100
gaggcaccgg gcacaggcaa ggagacagcc cccgaacggc gggagggtgt gacactgcgc 2160
tgcaaccacc gcgatgcaag tatgttgggc tcgctggcct acaatgtgct cctcatcgcg 2220
ctctgcacgc tttatgcctt caagactcgc aagtgcctcg aaaacttcaa cgaggccaag 2280
ttcattggct tcaccatgta caccacctgc atcatctggc tggcattcct gcccatcttc 2340
tatgtcacct ccagtgacta ccgggtacag accaccacca tgtgctgtgt agtcagcctc 2400
agcggctccg tgggtgcttg ctgcctcttt gcgcccagc tgcacatcat cctcttccag 2460
ccgcagaaga acaccatcga ggagggtgct tgcagcaccg cagctcacgc tttcaagggtg 2520
gctgcccggg ccacgctgcg ccgcagcaac gtctcccgca agcgggtccag cagccttgga 2580
ggctccacgg gatccacccc ctctctctcc atcagcagca agagcaacag cgaagaccca 2640
ttcccacagc ccgagaggca gaagcagcag cagccgctgg ccctaacca gcaagagcag 2700
cagcagcagc ccctgaccct cccacagcag caacgatctc agcagcagcc cagatgcaag 2760
cagaaggcca tctttggcag cggcacggct accttctcac tgagctttga tgagcctcag 2820
aagaacgcca tggcccacgg gaattctacg caccagaact ccctggaggc ccagaaaagc 2880
agcgatacgc tgacccgaca ccagccatta ctcccgtgc agtgccggga aacggactta 2940
gatctgaccg tccaggaaac aggtctgcaa ggacctgtgg gtggagacca gcggccagag 3000
gtggaggacc ctgaagagtt gtccccagca cttgtagtgt ccagttcaca gagctttgtc 3060
atcagtggtg gaggcagcac tgttacagaa aacgtagtga attca 3105

```

<210> 35

<211> 1035

<212> PRT

<213> Artificial Sequence

<220>

<223> Chimeric hmGluR2/hCaR

<400> 35

```

Met Gly Ser Leu Leu Ala Leu Pro Ala Leu Leu Leu Leu Trp Gly Ala
 1             5             10             15

```

```

Val Ala Glu Gly Pro Ala Lys Lys Val Leu Thr Leu Glu Gly Asp Leu
      20             25             30

```

```

Val Leu Gly Gly Leu Phe Pro Val His Gln Lys Gly Gly Pro Ala Glu
      35             40             45

```

```

Asp Cys Gly Pro Val Asn Glu His Arg Gly Ile Gln Arg Leu Glu Ala

```

50					55					60					
Met	Leu	Phe	Ala	Leu	Asp	Arg	Ile	Asn	Arg	Asp	Pro	His	Leu	Leu	Pro
65					70					75					80
Gly	Val	Arg	Leu	Gly	Ala	His	Ile	Leu	Asp	Ser	Cys	Ser	Lys	Asp	Thr
				85					90					95	
His	Ala	Leu	Glu	Gln	Ala	Leu	Asp	Phe	Val	Arg	Ala	Ser	Leu	Ser	Arg
			100					105					110		
Gly	Ala	Asp	Gly	Ser	Arg	His	Ile	Cys	Pro	Asp	Gly	Ser	Tyr	Ala	Thr
		115					120					125			
His	Gly	Asp	Ala	Pro	Thr	Ala	Ile	Thr	Gly	Val	Ile	Gly	Gly	Ser	Tyr
	130					135					140				
Ser	Asp	Val	Ser	Ile	Gln	Val	Ala	Asn	Leu	Leu	Arg	Leu	Phe	Gln	Ile
145					150					155					160
Pro	Gln	Ile	Ser	Tyr	Ala	Ser	Thr	Ser	Ala	Lys	Leu	Ser	Asp	Lys	Ser
				165					170					175	
Arg	Tyr	Asp	Tyr	Phe	Ala	Arg	Thr	Val	Pro	Pro	Asp	Phe	Phe	Gln	Ala
			180					185					190		
Lys	Ala	Met	Ala	Glu	Ile	Leu	Arg	Phe	Phe	Asn	Trp	Thr	Tyr	Val	Ser
	195						200					205			
Thr	Val	Ala	Ser	Glu	Gly	Asp	Tyr	Gly	Glu	Thr	Gly	Ile	Glu	Ala	Phe
	210					215					220				
Glu	Leu	Glu	Ala	Arg	Ala	Arg	Asn	Ile	Cys	Val	Ala	Thr	Ser	Glu	Lys
225					230					235				240	
Val	Gly	Arg	Ala	Met	Ser	Arg	Ala	Ala	Phe	Glu	Gly	Val	Val	Arg	Ala
				245					250					255	
Leu	Leu	Gln	Lys	Pro	Ser	Ala	Arg	Val	Ala	Val	Leu	Phe	Thr	Arg	Ser
			260					265					270		
Glu	Asp	Ala	Arg	Glu	Leu	Leu	Ala	Ala	Ser	Gln	Arg	Leu	Asn	Ala	Ser
	275						280					285			
Phe	Thr	Trp	Val	Ala	Ser	Asp	Gly	Trp	Gly	Ala	Leu	Glu	Ser	Val	Val
	290					295					300				
Ala	Gly	Ser	Glu	Gly	Ala	Ala	Glu	Gly	Ala	Ile	Thr	Ile	Glu	Leu	Ala
305					310					315				320	
Ser	Tyr	Pro	Ile	Ser	Asp	Phe	Ala	Ser	Tyr	Phe	Gln	Ser	Leu	Asp	Pro
				325					330					335	
Trp	Asn	Asn	Ser	Arg	Asn	Pro	Trp	Phe	Arg	Glu	Phe	Trp	Glu	Gln	Arg
			340					345					350		

Phe Arg Cys Ser Phe Arg Gln Arg Asp Cys Ala Ala His Ser Leu Arg  
 355 360 365  
 Ala Val Pro Phe Glu Gln Glu Ser Lys Ile Met Phe Val Val Asn Ala  
 370 375 380  
 Val Tyr Ala Met Ala His Ala Leu His Asn Met His Arg Ala Leu Cys  
 385 390 395 400  
 Pro Asn Thr Thr Arg Leu Cys Asp Ala Met Arg Pro Val Asn Gly Arg  
 405 410 415  
 Arg Leu Tyr Lys Asp Phe Val Leu Asn Val Lys Phe Asp Ala Pro Phe  
 420 425 430  
 Arg Pro Ala Asp Thr His Asn Glu Val Arg Phe Asp Arg Phe Gly Asp  
 435 440 445  
 Gly Ile Gly Arg Tyr Asn Ile Phe Thr Tyr Leu Arg Ala Gly Ser Gly  
 450 455 460  
 Arg Tyr Arg Tyr Gln Lys Val Gly Tyr Trp Ala Glu Gly Leu Thr Leu  
 465 470 475 480  
 Asp Thr Ser Leu Ile Pro Trp Ala Ser Pro Ser Ala Gly Pro Leu Pro  
 485 490 495  
 Ala Ser Arg Cys Ser Glu Pro Cys Leu Gln Asn Glu Val Lys Ser Val  
 500 505 510  
 Gln Pro Gly Glu Val Cys Cys Trp Leu Cys Ile Pro Cys Gln Pro Tyr  
 515 520 525  
 Glu Tyr Arg Leu Asp Glu Phe Thr Cys Ala Asp Cys Gly Leu Gly Tyr  
 530 535 540  
 Trp Pro Asn Ala Ser Leu Thr Gly Cys Phe Glu Leu Pro Gln Glu Tyr  
 545 550 555 560  
 Ile Arg Trp Gly Asp Ala Trp Ala Val Gly Pro Val Thr Ile Ala Cys  
 565 570 575  
 Leu Gly Ala Leu Ala Thr Leu Phe Val Leu Gly Val Phe Val Arg His  
 580 585 590  
 Asn Ala Thr Pro Val Val Lys Ala Ser Gly Arg Glu Leu Cys Tyr Ile  
 595 600 605  
 Leu Leu Gly Gly Val Phe Leu Cys Tyr Cys Met Thr Phe Ile Phe Ile  
 610 615 620  
 Ala Lys Pro Ser Thr Ala Val Cys Thr Leu Arg Arg Leu Gly Leu Gly  
 625 630 635 640  
 Thr Ala Phe Ser Val Cys Tyr Ser Ala Leu Leu Thr Lys Thr Asn Arg

			645				650				655				
Ile	Ala	Arg	Ile 660	Phe	Gly	Gly	Ala	Arg	Glu	Gly	Ala	Gln	Arg	Pro	Arg
Phe	Ile	Ser	Pro 675	Ala	Ser	Gln	Val	Ala	Ile	Cys	Leu	Ala	Leu	Ile	Ser
Gly	Gln	Leu	Leu 690	Ile	Val	Val	Ala	Trp	Leu	Val	Val	Glu	Ala	Pro	Gly
Thr 705	Gly	Lys	Glu	Thr	Ala 710	Pro	Glu	Arg	Arg	Glu 715	Val	Val	Thr	Leu	Arg 720
Cys	Asn	His	Arg 725	Asp	Ala	Ser	Met	Leu	Gly 730	Ser	Leu	Ala	Tyr	Asn 735	Val
Leu	Leu	Ile	Ala 740	Leu	Cys	Thr	Leu	Tyr	Ala 745	Phe	Lys	Thr	Arg 750	Lys	Cys
Pro	Glu	Asn	Phe 755	Asn	Glu	Ala	Lys	Phe	Ile	Gly	Phe	Thr	Met 765	Tyr	Thr
Thr 770	Cys	Ile	Ile	Trp	Leu	Ala 775	Phe	Leu	Pro	Ile	Phe 780	Tyr	Val	Thr	Ser
Ser 785	Asp	Tyr	Arg	Val	Gln 790	Thr	Thr	Thr	Met	Cys 795	Val	Ser	Val	Ser	Leu 800
Ser	Gly	Ser	Val 805	Val	Leu	Gly	Cys	Leu	Phe 810	Ala	Pro	Lys	Leu	His 815	Ile
Ile	Leu	Phe	Gln 820	Pro	Gln	Lys	Asn	Thr 825	Ile	Glu	Glu	Val	Arg 830	Cys	Ser
Thr	Ala	Ala	His 835	Ala	Phe	Lys	Val 840	Ala	Ala	Arg	Ala	Thr 845	Leu	Arg	Arg
Ser 850	Asn	Val	Ser	Arg	Lys	Arg 855	Ser	Ser	Ser	Leu	Gly 860	Gly	Ser	Thr	Gly
Ser 865	Thr	Pro	Ser	Ser	Ser 870	Ile	Ser	Ser	Lys	Ser 875	Asn	Ser	Glu	Asp	Pro 880
Phe	Pro	Gln	Pro	Glu 885	Arg	Gln	Lys	Gln	Gln 890	Gln	Pro	Leu	Ala	Leu 895	Thr
Gln	Gln	Glu	Gln 900	Gln	Gln	Gln	Pro	Leu 905	Thr	Leu	Pro	Gln	Gln 910	Gln	Arg
Ser	Gln	Gln	Gln 915	Pro	Arg	Cys	Lys 920	Gln	Lys	Val	Ile	Phe 925	Gly	Ser	Gly
Thr 930	Val	Thr	Phe	Ser	Leu	Ser 935	Phe	Asp	Glu	Pro	Gln 940	Lys	Asn	Ala	Met

Ala His Gly Asn Ser Thr His Gln Asn Ser Leu Glu Ala Gln Lys Ser  
 945 950 955 960

Ser Asp Thr Leu Thr Arg His Gln Pro Leu Leu Pro Leu Gln Cys Gly  
 965 970 975

Glu Thr Asp Leu Asp Leu Thr Val Gln Glu Thr Gly Leu Gln Gly Pro  
 980 985 990

Val Gly Gly Asp Gln Arg Pro Glu Val Glu Asp Pro Glu Glu Leu Ser  
 995 1000 1005

Pro Ala Leu Val Val Ser Ser Ser Gln Ser Phe Val Ile Ser Gly Gly  
 1010 1015 1020

Gly Ser Thr Val Thr Glu Asn Val Val Asn Ser  
 1025 1030 1035

<210> 36  
 <211> 4185  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Chimeric pmGluR2//CaR\*G(qi5

<400> 36

atgggatcgc	tgcttgcgct	cccggcactg	ctgctgctgt	ggggtgctgt	ggctgagggc	60
ccagccaaga	aggtgctgac	cctggagggg	gacttggtgc	tgggtgggct	gttcccagtg	120
caccagaagg	gcgggccagc	agaggactgt	ggtcctgtca	atgagcaccg	tggcatccag	180
cgcttgagg	ccatgctttt	tgcactggac	cgcacacacc	gtgacccgca	cctgctgcct	240
ggcgtgcgcc	tgggtgcaca	catcctcgac	agttgctcca	aggacacaca	tgcgctggag	300
caggcactgg	actttgtgcg	tgcctcactc	agcctggtg	ctgatggctc	acgccacatc	360
tgccccgacg	gctcttatgc	gacccatggt	gatgctccca	ctgccatcac	tgggtgtatt	420
ggcggttcct	acagtgatgt	ctccatccag	gtggccaacc	tcttgaggct	atttcagatc	480
ccacagatta	gctacgcctc	taccagtgcc	aagctgagtg	acaagtcccg	ctatgactac	540
tttggccgca	cagtgcctcc	tgacttcttc	caagccaagg	ccatggctga	gattctccgc	600
ttcttcaact	ggacctatgt	gtccactgtg	gcgtctgagg	gcgactatgg	cgagacaggc	660
attgaggcct	ttgagctaga	ggctcgtgcc	cgcaacatct	gtgtggccac	ctcggagaaa	720
gtgggcccgtg	ccatgagccg	cgcggccctt	gaggggtgtg	tgcgagccct	gctgcagaag	780
cccagtgcc	gcgtggctgt	cctgttcacc	cgttctgagg	atgcccgga	gctgcttgct	840
gccagccagc	gcctcaatgc	cagcttcacc	tgggtggcca	gtgatggtg	gggggccctg	900
gagagtgtgg	tggcaggcag	tgagggggct	gctgaggggt	ctatcaccat	cgagctggcc	960
tcctacccca	tcagtgactt	tgcctcctac	ttccagagcc	tggacccttg	gaacaacagc	1020
cggaaccctt	ggttccgtga	attctgggag	cagaggttcc	gctgcagctt	ccggcagcga	1080
gactgcgcag	cccactctct	ccgggctgtg	ccctttgagc	aggagtccaa	gatcatgttt	1140
gtggtcaatg	cagtgtacgc	catggcccat	gcgctccaca	acatgcaccg	tgccctctgc	1200
cccaacacca	cccggctctg	tgacgcgatg	cggccagtta	acgggcgcgc	cctctacaag	1260
gactttgtgc	tcaacgtcaa	gtttgatgcc	ccctttcgcc	cagctgacac	ccacaatgag	1320
gtccgctttg	accgctttgg	tgatggtatt	ggccgctaca	acatcttcac	ctatctgcgt	1380
gcaggcagtg	ggcgctatcg	ctaccagaag	gtgggctact	gggcagaagg	cttgactctg	1440
gacaccagcc	tcaccccatg	ggcctcacc	tcagccggcc	ccctgcccgc	ctctcgctgc	1500
agtgagccct	gcctccagaa	tgaggtgaag	agtgtgcagc	cgggcgaagt	ctgctgctgg	1560

```

ctctgcattc cgtgccagcc ctatgagtac cgattggacg aattcacttg cgctgattgt 1620
ggcctggggt actggcccaa tgccagcctg actggctgct tcgaactgcc ccaggagtac 1680
atccgctggg gcgatgcctg ggctgtggga cctgtcacca tcgctgcct cggtgccctg 1740
gccacctctt ttgtgctggg tgtctttgtg cggcacaaat ccacaccagt ggtcaaggcc 1800
tcaggteggy agctctgcta catcctgctg ggtgggtgtc tcctctgcta ctgcatgacc 1860
ttcatcttca ttgccaagcc atccacggca gtgtgtacct tacggcgtct tggtttgggc 1920
actgccttct ctgtctgcta ctcagccctg ctaccaaga ccaaccgcat tgcacgcac 1980
ttcgggtggg cccgggaggg tgcccagcgg ccacgcttca tcagtccctg ctcacagggtg 2040
gccatctgcc tggcacttat ctcgggccag ctgctcatcg tggtcgcctg gctgggtgggtg 2100
gaggcaccgg gcacaggcaa ggagacagcc cccgaacggc gggagggtgg gacactgcgc 2160
tgcaaccacc gcgatgcaag tatgttgggc tcgctggcct acaatgtgct cctcatcgcg 2220
ctctgcacgc tttatgcctt caagactcgc aagtgcctcg aaaacttcaa cgaggccaag 2280
ttcattgggt tcacatgta caccacctgc atcatctggc tggcattcct gcccatcttc 2340
tatgtcacct ccagtgaacta cgggtacag accaccacca tgtgcgtgtc agtcagcctc 2400
agcggtccg tgggtgcttg ctgcctcttt gcgccaagc tgcacatcat cctcttccag 2460
ccgcagaaga acaccatcga ggagggtcgt tgcagcaccg cagctcacgc tttcaagggtg 2520
gctgcccggg ccacgctgcg ccgcagcaac gtctcccgca agcggtccag cagccttggg 2580
ggctccacgg gatccacccc ctctctctcc atcagcagca agagcaacag cgaagaccca 2640
ttcccacagc ccgagaggca gaagcagcag cagccgctgg ccctaacca gcaagagcag 2700
cagcagcagc ccctgaccct cccacagcag caacgatctc agcagcagcc cagatgcaag 2760
cagaagggtca tctttggcag cggcacgggtc accttctcac tgagctttga tgagcctcag 2820
aagaacgcca tggcccacgg gaattctacg caccagaact ccctggaggc ccagaaaagc 2880
agcgatacgc tgaccgcaga ccagccatta ctcccgtgc agtgcgggga aacggactta 2940
gatctgaccg tccaggaaac aggtctgcaa ggacctgtgg gtggagacca gcggccagag 3000
gtggaggacc ctgaagagtt gtccccagca cttgtagtgt ccagttcaca gagctttgtc 3060
atcagtgggt gaggcagcac tgttacagaa aacgtagtga attcaatgac tctggagtcc 3120
atcatggcgt gctgcctgag cgaggaggcc aagggaagccc ggcggatcaa cgacgagatc 3180
gagcggcagc tccgcaggga caagcgggac gccgcgggg agtcaagct gctgctgctc 3240
gggacaggag agagtggcaa gagtacgttt atcaagcaga tgagaatcat ccatgggtca 3300
ggatactctg atgaagataa aaggggcttc accaagctgg tgtatcagaa catcttcacg 3360
gccatgcagc ccatgatcag agccatggac aactcaaga tcccatataa gtatgagcac 3420
aataaggctc atgcacaatt agttcgagaa gttgatgtgg agaagggtgtc tgcttttgag 3480
aatccatatg tagatgcaat aaagagttta tggaaatgat ctggaatcca ggaatgctat 3540
gatagacgac gagaatatca attatctgac tctaccaaact actatcttaa tgacttgagc 3600
cgcgtagctg accctgccta cctgcctacg caacaagatg tgcttagagt tcgagtcccc 3660
accacaggga tcatcgaata cccctttgac ttacaaagtg tcattttcag aatggtcgat 3720
gtagggggcc aaaggctcaga gagaagaaaa tggatacact gctttgaaaa tgtcacctct 3780
atcatgtttc tagtagcgct tagtgaatat gatcaagttc tcgtggagtc agacaatgag 3840
aaccgaatgg aggaaagcaa ggctctcttt agaacaatta tcacataccc ctggttccag 3900
aactcctcgg ttattctgtt cttaaacaaag aaagatcttc tagaggagaa aatcatgtat 3960
tcccatctag tcgactactt cccagaatat gatggacccc agagagatgc ccaggcagcc 4020
cgagaattca ttctgaagat gttcgtggac ctgaaccag acagtgcaca aattatctac 4080
tcccacttca cgtgcgccac agacaccgag aatatccgct ttgtctttgc tgccgtcaag 4140
gacaccatcc tccagttgaa cctgaaggac tgcggctctgt tctaa 4185

```

<210> 37

<211> 1394

<212> PRT

<213> Artificial Sequence

<220>

<223> Chimeric pmGluR2//CaR\*G(qi5

&lt;400&gt; 37

Met	Gly	Ser	Leu	Leu	Ala	Leu	Pro	Ala	Leu	Leu	Leu	Leu	Trp	Gly	Ala	1	5	10	15
Val	Ala	Glu	Gly	Pro	Ala	Lys	Lys	Val	Leu	Thr	Leu	Glu	Gly	Asp	Leu	20	25	30	
Val	Leu	Gly	Gly	Leu	Phe	Pro	Val	His	Gln	Lys	Gly	Gly	Pro	Ala	Glu	35	40	45	
Asp	Cys	Gly	Pro	Val	Asn	Glu	His	Arg	Gly	Ile	Gln	Arg	Leu	Glu	Ala	50	55	60	
Met	Leu	Phe	Ala	Leu	Asp	Arg	Ile	Asn	Arg	Asp	Pro	His	Leu	Leu	Pro	65	70	75	80
Gly	Val	Arg	Leu	Gly	Ala	His	Ile	Leu	Asp	Ser	Cys	Ser	Lys	Asp	Thr	85	90	95	
His	Ala	Leu	Glu	Gln	Ala	Leu	Asp	Phe	Val	Arg	Ala	Ser	Leu	Ser	Arg	100	105	110	
Gly	Ala	Asp	Gly	Ser	Arg	His	Ile	Cys	Pro	Asp	Gly	Ser	Tyr	Ala	Thr	115	120	125	
His	Gly	Asp	Ala	Pro	Thr	Ala	Ile	Thr	Gly	Val	Ile	Gly	Gly	Ser	Tyr	130	135	140	
Ser	Asp	Val	Ser	Ile	Gln	Val	Ala	Asn	Leu	Leu	Arg	Leu	Phe	Gln	Ile	145	150	155	160
Pro	Gln	Ile	Ser	Tyr	Ala	Ser	Thr	Ser	Ala	Lys	Leu	Ser	Asp	Lys	Ser	165	170	175	
Arg	Tyr	Asp	Tyr	Phe	Ala	Arg	Thr	Val	Pro	Pro	Asp	Phe	Phe	Gln	Ala	180	185	190	
Lys	Ala	Met	Ala	Glu	Ile	Leu	Arg	Phe	Phe	Asn	Trp	Thr	Tyr	Val	Ser	195	200	205	
Thr	Val	Ala	Ser	Glu	Gly	Asp	Tyr	Gly	Glu	Thr	Gly	Ile	Glu	Ala	Phe	210	215	220	
Glu	Leu	Glu	Ala	Arg	Ala	Arg	Asn	Ile	Cys	Val	Ala	Thr	Ser	Glu	Lys	225	230	235	240
Val	Gly	Arg	Ala	Met	Ser	Arg	Ala	Ala	Phe	Glu	Gly	Val	Val	Arg	Ala	245	250	255	
Leu	Leu	Gln	Lys	Pro	Ser	Ala	Arg	Val	Ala	Val	Leu	Phe	Thr	Arg	Ser	260	265	270	
Glu	Asp	Ala	Arg	Glu	Leu	Leu	Ala	Ala	Ser	Gln	Arg	Leu	Asn	Ala	Ser	275	280	285	



Phe Thr Trp Val Ala Ser Asp Gly Trp Gly Ala Leu Glu Ser Val Val  
 290 295 300  
 Ala Gly Ser Glu Gly Ala Ala Glu Gly Ala Ile Thr Ile Glu Leu Ala  
 305 310 315 320  
 Ser Tyr Pro Ile Ser Asp Phe Ala Ser Tyr Phe Gln Ser Leu Asp Pro  
 325 330 335  
 Trp Asn Asn Ser Arg Asn Pro Trp Phe Arg Glu Phe Trp Glu Gln Arg  
 340 345 350  
 Phe Arg Cys Ser Phe Arg Gln Arg Asp Cys Ala Ala His Ser Leu Arg  
 355 360 365  
 Ala Val Pro Phe Glu Gln Glu Ser Lys Ile Met Phe Val Val Asn Ala  
 370 375 380  
 Val Tyr Ala Met Ala His Ala Leu His Asn Met His Arg Ala Leu Cys  
 385 390 395 400  
 Pro Asn Thr Thr Arg Leu Cys Asp Ala Met Arg Pro Val Asn Gly Arg  
 405 410 415  
 Arg Leu Tyr Lys Asp Phe Val Leu Asn Val Lys Phe Asp Ala Pro Phe  
 420 425 430  
 Arg Pro Ala Asp Thr His Asn Glu Val Arg Phe Asp Arg Phe Gly Asp  
 435 440 445  
 Gly Ile Gly Arg Tyr Asn Ile Phe Thr Tyr Leu Arg Ala Gly Ser Gly  
 450 455 460  
 Arg Tyr Arg Tyr Gln Lys Val Gly Tyr Trp Ala Glu Gly Leu Thr Leu  
 465 470 475 480  
 Asp Thr Ser Leu Ile Pro Trp Ala Ser Pro Ser Ala Gly Pro Leu Pro  
 485 490 495  
 Ala Ser Arg Cys Ser Glu Pro Cys Leu Gln Asn Glu Val Lys Ser Val  
 500 505 510  
 Gln Pro Gly Glu Val Cys Cys Trp Leu Cys Ile Pro Cys Gln Pro Tyr  
 515 520 525  
 Glu Tyr Arg Leu Asp Glu Phe Thr Cys Ala Asp Cys Gly Leu Gly Tyr  
 530 535 540  
 Trp Pro Asn Ala Ser Leu Thr Gly Cys Phe Glu Leu Pro Gln Glu Tyr  
 545 550 555 560  
 Ile Arg Trp Gly Asp Ala Trp Ala Val Gly Pro Val Thr Ile Ala Cys  
 565 570 575  
 Leu Gly Ala Leu Ala Thr Leu Phe Val Leu Gly Val Phe Val Arg His  
 580 585 590

Asn Ala Thr Pro Val Val Lys Ala Ser Gly Arg Glu Leu Cys Tyr Ile  
 595 600 605  
 Leu Leu Gly Gly Val Phe Leu Cys Tyr Cys Met Thr Phe Ile Phe Ile  
 610 615 620  
 Ala Lys Pro Ser Thr Ala Val Cys Thr Leu Arg Arg Leu Gly Leu Gly  
 625 630 635 640  
 Thr Ala Phe Ser Val Cys Tyr Ser Ala Leu Leu Thr Lys Thr Asn Arg  
 645 650 655  
 Ile Ala Arg Ile Phe Gly Gly Ala Arg Glu Gly Ala Gln Arg Pro Arg  
 660 665 670  
 Phe Ile Ser Pro Ala Ser Gln Val Ala Ile Cys Leu Ala Leu Ile Ser  
 675 680 685  
 Gly Gln Leu Leu Ile Val Val Ala Trp Leu Val Val Glu Ala Pro Gly  
 690 695 700  
 Thr Gly Lys Glu Thr Ala Pro Glu Arg Arg Glu Val Val Thr Leu Arg  
 705 710 715 720  
 Cys Asn His Arg Asp Ala Ser Met Leu Gly Ser Leu Ala Tyr Asn Val  
 725 730 735  
 Leu Leu Ile Ala Leu Cys Thr Leu Tyr Ala Phe Lys Thr Arg Lys Cys  
 740 745 750  
 Pro Glu Asn Phe Asn Glu Ala Lys Phe Ile Gly Phe Thr Met Tyr Thr  
 755 760 765  
 Thr Cys Ile Ile Trp Leu Ala Phe Leu Pro Ile Phe Tyr Val Thr Ser  
 770 775 780  
 Ser Asp Tyr Arg Val Gln Thr Thr Thr Met Cys Val Ser Val Ser Leu  
 785 790 795 800  
 Ser Gly Ser Val Val Leu Gly Cys Leu Phe Ala Pro Lys Leu His Ile  
 805 810 815  
 Ile Leu Phe Gln Pro Gln Lys Asn Thr Ile Glu Glu Val Arg Cys Ser  
 820 825 830  
 Thr Ala Ala His Ala Phe Lys Val Ala Ala Arg Ala Thr Leu Arg Arg  
 835 840 845  
 Ser Asn Val Ser Arg Lys Arg Ser Ser Ser Leu Gly Gly Ser Thr Gly  
 850 855 860  
 Ser Thr Pro Ser Ser Ser Ile Ser Ser Lys Ser Asn Ser Glu Asp Pro  
 865 870 875 880  
 Phe Pro Gln Pro Glu Arg Gln Lys Gln Gln Gln Pro Leu Ala Leu Thr

885										890					895					
Gln	Gln	Glu	Gln	Gln	Gln	Gln	Pro	Leu	Thr	Leu	Pro	Gln	Gln	Gln	Arg					
900										905					910					
Ser	Gln	Gln	Gln	Pro	Arg	Cys	Lys	Gln	Lys	Val	Ile	Phe	Gly	Ser	Gly					
915										920					925					
Thr	Val	Thr	Phe	Ser	Leu	Ser	Phe	Asp	Glu	Pro	Gln	Lys	Asn	Ala	Met					
930										935					940					
Ala	His	Gly	Asn	Ser	Thr	His	Gln	Asn	Ser	Leu	Glu	Ala	Gln	Lys	Ser					
945										950					955					
Ser	Asp	Thr	Leu	Thr	Arg	His	Gln	Pro	Leu	Leu	Pro	Leu	Gln	Cys	Gly					
965										970					975					
Glu	Thr	Asp	Leu	Asp	Leu	Thr	Val	Gln	Glu	Thr	Gly	Leu	Gln	Gly	Pro					
980										985					990					
Val	Gly	Gly	Asp	Gln	Arg	Pro	Glu	Val	Glu	Asp	Pro	Glu	Glu	Leu	Ser					
995										1000					1005					
Pro	Ala	Leu	Val	Val	Ser	Ser	Ser	Gln	Ser	Phe	Val	Ile	Ser	Gly	Gly					
1010										1015					1020					
Gly	Ser	Thr	Val	Thr	Glu	Asn	Val	Val	Asn	Ser	Met	Thr	Leu	Glu	Ser					
1025										1030					1035					
Ile	Met	Ala	Cys	Cys	Leu	Ser	Glu	Glu	Ala	Lys	Glu	Ala	Arg	Arg	Ile					
1045										1050					1055					
Asn	Asp	Glu	Ile	Glu	Arg	Gln	Leu	Arg	Arg	Asp	Lys	Arg	Asp	Ala	Arg					
1060										1065					1070					
Arg	Glu	Leu	Lys	Leu	Leu	Leu	Leu	Gly	Thr	Gly	Glu	Ser	Gly	Lys	Ser					
1075										1080					1085					
Thr	Phe	Ile	Lys	Gln	Met	Arg	Ile	Ile	His	Gly	Ser	Gly	Tyr	Ser	Asp					
1090										1095					1100					
Glu	Asp	Lys	Arg	Gly	Phe	Thr	Lys	Leu	Val	Tyr	Gln	Asn	Ile	Phe	Thr					
1105										1110					1115					
Ala	Met	Gln	Ala	Met	Ile	Arg	Ala	Met	Asp	Thr	Leu	Lys	Ile	Pro	Tyr					
1125										1130					1135					
Lys	Tyr	Glu	His	Asn	Lys	Ala	His	Ala	Gln	Leu	Val	Arg	Glu	Val	Asp					
1140										1145					1150					
Val	Glu	Lys	Val	Ser	Ala	Phe	Glu	Asn	Pro	Tyr	Val	Asp	Ala	Ile	Lys					
1155										1160					1165					
Ser	Leu	Trp	Asn	Asp	Pro	Gly	Ile	Gln	Glu	Cys	Tyr	Asp	Arg	Arg	Arg					
1170										1175					1180					

Glu Tyr Gln Leu Ser Asp Ser Thr Lys Tyr Tyr Leu Asn Asp Leu Asp  
1185 1190 1195 1200

Arg Val Ala Asp Pro Ala Tyr Leu Pro Thr Gln Gln Asp Val Leu Arg  
1205 1210 1215

Val Arg Val Pro Thr Thr Gly Ile Ile Glu Tyr Pro Phe Asp Leu Gln  
1220 1225 1230

Ser Val Ile Phe Arg Met Val Asp Val Gly Gly Gln Arg Ser Glu Arg  
1235 1240 1245

Arg Lys Trp Ile His Cys Phe Glu Asn Val Thr Ser Ile Met Phe Leu  
1250 1255 1260

Val Ala Leu Ser Glu Tyr Asp Gln Val Leu Val Glu Ser Asp Asn Glu  
1265 1270 1275 1280

Asn Arg Met Glu Glu Ser Lys Ala Leu Phe Arg Thr Ile Ile Thr Tyr  
1285 1290 1295

Pro Trp Phe Gln Asn Ser Ser Val Ile Leu Phe Leu Asn Lys Lys Asp  
1300 1305 1310

Leu Leu Glu Glu Lys Ile Met Tyr Ser His Leu Val Asp Tyr Phe Pro  
1315 1320 1325

Glu Tyr Asp Gly Pro Gln Arg Asp Ala Gln Ala Ala Arg Glu Phe Ile  
1330 1335 1340

Leu Lys Met Phe Val Asp Leu Asn Pro Asp Ser Asp Lys Ile Ile Tyr  
1345 1350 1355 1360

Ser His Phe Thr Cys Ala Thr Asp Thr Glu Asn Ile Arg Phe Val Phe  
1365 1370 1375

Ala Ala Val Lys Asp Thr Ile Leu Gln Leu Asn Leu Lys Asp Cys Gly  
1380 1385 1390

Leu Phe

<210> 38

<211> 3177

<212> DNA

<213> Artificial Sequence

<220>

<223> Chimeric hmGluR8/hCaR

<400> 38

atggtatgcg agggaaagcg atcagcctct tgccttggt tcttctctt gaccgccaag  
ttctactgga tcctcacaat gatgcaaaga actcacagcc aggagtatgc ccattccata

60

120

cggggtggatg	gggacattat	tttggggggg	ctcttccctg	tccacgcaaa	gggagagaga	180
gggggtgcctt	gtggggagct	gaagaaggaa	aaggggattc	acagactgga	ggccatgctt	240
tatgcaattg	accagattaa	caaggaccct	gatctccttt	ccaacatcac	tctgggtgtc	300
cgcatacctcg	acacgtgctc	tagggacacc	tatgcttttg	agcagtctct	aacattcgtg	360
caggcattaa	tagagaaaga	tgcttcggat	gtgaagtgtg	ctaattggaga	tccaccatt	420
ttcaccaagc	ccgacaagat	ttctggcgctc	atagggtgctg	cagcaagctc	cgtgtccatc	480
atggttgcta	acattttaag	actttttaag	atacctcaaa	tcagctatgc	atccacagcc	540
ccagagctaa	gtgataacac	cagggtatgac	tttttctctc	gagtggttcc	gcctgactcc	600
taccaagccc	aagccatggt	ggacatcgctg	acagcactgg	gatggaatta	tgtttcgaca	660
ctggcttctg	aggggaacta	tggtgagagc	ggtgtggagg	ccttcaccca	gatctcgagg	720
gagattggtg	gtgtttgcat	tgctcagtc	cagaaaatcc	cacgtgaacc	aagacctgga	780
gaatttgaaa	aaattatcaa	acgcctgcta	gaaacaccta	atgctcgagc	agtgattatg	840
tttgccaatg	aggatgacat	caggaggata	ttggaagcag	caaaaaaact	aaaccaaaagt	900
gggcattttc	tctggatttg	ctcagatagt	tggggatcca	aaatagcacc	tgtctatcag	960
caagaggaga	ttgcagaagg	ggctgtgaca	attttgccca	aacgagcatc	aattgatgga	1020
tttgatcgat	actttagaag	ccgaactctt	gccaataatc	gaagaaatgt	gtggtttgca	1080
gaattctggg	aggagaat	tggtcgcaag	ttaggatcac	atgggaaaag	gaacagtc	1140
ataaagaaat	gcacagggct	ggagcgaatt	gctcgggatt	catcttatga	acaggaagga	1200
aaggtccaat	ttgtaattga	tgctgtatat	tccatggctt	acgccctgca	caatatgcac	1260
aaagatctct	gocctggata	cattggcctt	tgtccacgaa	tgagtaccat	tgatgggaaa	1320
gagctacttg	gttatattcg	ggctgtaaat	tttaatggca	gtgctggcac	tctgtcact	1380
tttaatgaaa	acggagatgc	tcttgacgt	tatgatatct	tccagtatca	aataaccaac	1440
aaaagcacag	agtacaaagt	catcgccac	tggaccaatc	agcttcatct	aaaagtggaa	1500
gacatgcagt	gggctcatag	agaacatact	cacccggcgt	ctgtctgcag	cctgccgtgt	1560
aagccagggg	agaggaagaa	aacggtgaaa	gggggtccctt	gctgctggca	ctgtgaacgc	1620
tgtgaagggt	acaactacca	ggtggatgag	ctgtcctgtg	aactttgccc	tctggatcag	1680
agaccaaca	tgaaccgcac	aggctgccag	cttatcccca	tcatacaaatt	ggagtggcat	1740
tctccctggg	ctgtggtgcc	tgtgtttggt	gcaatattgg	gaatcatcgc	caccactttt	1800
gtgatcgtga	cctttgtccg	ctataatgac	acacctaactg	tgagggtctc	aggacgcgaa	1860
cttagttacg	tgctcctaac	ggggattttt	ctctgttatt	caatcacgtt	tttaatgatt	1920
gcagcaccag	atacaatcat	atgctccttc	cgacgggtct	tcctaggact	tggcatgtgt	1980
ttcagctatg	gacccttct	gacaaaaca	aaccgtatcc	accgaatatt	tgagcagggg	2040
aagaaatctg	tcacagcgcc	caagttcatt	agtccagcat	ctcagctggt	gatcaccttc	2100
agcctcatct	ccgtccagct	ccttgagtg	tttgtctggt	ttgttgtgga	tccccccac	2160
atcatcattg	actatggaga	gcagcggaca	ctagatccag	agaaggccag	gggagtgtc	2220
aagtgtgaca	tttctgatct	ctcaactcatt	tgttcacttg	gatacagtat	cctcttgatg	2280
gtcacttgta	ctgtttatgc	cattaaaacg	agaggtgtcc	cagagacttt	caatgaagcc	2340
aaacctattg	gatttaccat	gtataccacc	tgcatacttt	ggttagcttt	catccccatc	2400
tttttttgta	cagcccagtc	agcagaaaag	atgtacatcc	agacaacaac	acttactgtc	2460
tccatgagtt	taagtgtctc	agtatctctg	ggcatgctct	atatgcccaa	ggtttatatt	2520
ataatttttc	atccagaaca	gaataccatc	gaggaggtgc	gttgacagcac	cgcagctcac	2580
gctttcaagg	tggctgcccg	ggccacgctg	cgccgcagca	acgtctcccg	caagcggctc	2640
agcagccttg	gaggctccac	gggatccacc	ccctcctcct	ccatcagcag	caagagcaac	2700
agcgaagacc	cattcccaca	gcccagagag	cagaagcagc	agcagccgct	ggccctaacc	2760
cagcaagagc	agcagcagca	gcccctgacc	ctcccacagc	agcaacgata	tcagcagcag	2820
cccagatgca	agcagaaggt	catctttggc	agcggcacgg	tcaccttctc	actgagcttt	2880
gatgagcctc	agaagaacgc	catggcccac	gggaattcta	cgcaccagaa	ctccctggag	2940
gccagaaaa	gcagcgatac	gctgaccgga	caccagccat	tactcccgtc	gcagtgcggg	3000
gaaacggact	tagatctgac	cgtccaggaa	atggtctctg	aaggacctgt	gggtggagac	3060
cagcggccag	aggtggagga	ccctgaagag	ttgtccccag	cacttgtagt	gtccagttca	3120
cagagctttg	tcatacagtg	tggaggcagc	actgttacag	aaaacgtagt	gaattca	3177

<211> 1059  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Chimeric hmGluR8/hCaR

<400> 39

Met	Val	Cys	Glu	Gly	Lys	Arg	Ser	Ala	Ser	Cys	Pro	Cys	Phe	Phe	Leu	1	5	10	15
Leu	Thr	Ala	Lys	Phe	Tyr	Trp	Ile	Leu	Thr	Met	Met	Gln	Arg	Thr	His	20	25	30	
Ser	Gln	Glu	Tyr	Ala	His	Ser	Ile	Arg	Val	Asp	Gly	Asp	Ile	Ile	Leu	35	40	45	
Gly	Gly	Leu	Phe	Pro	Val	His	Ala	Lys	Gly	Glu	Arg	Gly	Val	Pro	Cys	50	55	60	
Gly	Glu	Leu	Lys	Lys	Glu	Lys	Gly	Ile	His	Arg	Leu	Glu	Ala	Met	Leu	65	70	75	
Tyr	Ala	Ile	Asp	Gln	Ile	Asn	Lys	Asp	Pro	Asp	Leu	Leu	Ser	Asn	Ile	85	90	95	
Thr	Leu	Gly	Val	Arg	Ile	Leu	Asp	Thr	Cys	Ser	Arg	Asp	Thr	Tyr	Ala	100	105	110	
Leu	Glu	Gln	Ser	Leu	Thr	Phe	Val	Gln	Ala	Leu	Ile	Glu	Lys	Asp	Ala	115	120	125	
Ser	Asp	Val	Lys	Cys	Ala	Asn	Gly	Asp	Pro	Pro	Ile	Phe	Thr	Lys	Pro	130	135	140	
Asp	Lys	Ile	Ser	Gly	Val	Ile	Gly	Ala	Ala	Ala	Ser	Ser	Val	Ser	Ile	145	150	155	
Met	Val	Ala	Asn	Ile	Leu	Arg	Leu	Phe	Lys	Ile	Pro	Gln	Ile	Ser	Tyr	165	170	175	
Ala	Ser	Thr	Ala	Pro	Glu	Leu	Ser	Asp	Asn	Thr	Arg	Tyr	Asp	Phe	Phe	180	185	190	
Ser	Arg	Val	Val	Pro	Pro	Asp	Ser	Tyr	Gln	Ala	Gln	Ala	Met	Val	Asp	195	200	205	
Ile	Val	Thr	Ala	Leu	Gly	Trp	Asn	Tyr	Val	Ser	Thr	Leu	Ala	Ser	Glu	210	215	220	
Gly	Asn	Tyr	Gly	Glu	Ser	Gly	Val	Glu	Ala	Phe	Thr	Gln	Ile	Ser	Arg	225	230	235	
Glu	Ile	Gly	Gly	Val	Cys	Ile	Ala	Gln	Ser	Gln	Lys	Ile	Pro	Arg	Glu	245	250	255	

1059  
 PRT  
 Artificial Sequence  
 Chimeric hmGluR8/hCaR  
 39

Pro Arg Pro Gly Glu Phe Glu Lys Ile Ile Lys Arg Leu Leu Glu Thr  
 260 265 270  
 Pro Asn Ala Arg Ala Val Ile Met Phe Ala Asn Glu Asp Asp Ile Arg  
 275 280 285  
 Arg Ile Leu Glu Ala Ala Lys Lys Leu Asn Gln Ser Gly His Phe Leu  
 290 295 300  
 Trp Ile Gly Ser Asp Ser Trp Gly Ser Lys Ile Ala Pro Val Tyr Gln  
 305 310 315 320  
 Gln Glu Glu Ile Ala Glu Gly Ala Val Thr Ile Leu Pro Lys Arg Ala  
 325 330 335  
 Ser Ile Asp Gly Phe Asp Arg Tyr Phe Arg Ser Arg Thr Leu Ala Asn  
 340 345 350  
 Asn Arg Arg Asn Val Trp Phe Ala Glu Phe Trp Glu Glu Asn Phe Gly  
 355 360 365  
 Cys Lys Leu Gly Ser His Gly Lys Arg Asn Ser His Ile Lys Lys Cys  
 370 375 380  
 Thr Gly Leu Glu Arg Ile Ala Arg Asp Ser Ser Tyr Glu Gln Glu Gly  
 385 390 395 400  
 Lys Val Gln Phe Val Ile Asp Ala Val Tyr Ser Met Ala Tyr Ala Leu  
 405 410 415  
 His Asn Met His Lys Asp Leu Cys Pro Gly Tyr Ile Gly Leu Cys Pro  
 420 425 430  
 Arg Met Ser Thr Ile Asp Gly Lys Glu Leu Leu Gly Tyr Ile Arg Ala  
 435 440 445  
 Val Asn Phe Asn Gly Ser Ala Gly Thr Pro Val Thr Phe Asn Glu Asn  
 450 455 460  
 Gly Asp Ala Pro Gly Arg Tyr Asp Ile Phe Gln Tyr Gln Ile Thr Asn  
 465 470 475 480  
 Lys Ser Thr Glu Tyr Lys Val Ile Gly His Trp Thr Asn Gln Leu His  
 485 490 495  
 Leu Lys Val Glu Asp Met Gln Trp Ala His Arg Glu His Thr His Pro  
 500 505 510  
 Ala Ser Val Cys Ser Leu Pro Cys Lys Pro Gly Glu Arg Lys Lys Thr  
 515 520 525  
 Val Lys Gly Val Pro Cys Cys Trp His Cys Glu Arg Cys Glu Gly Tyr  
 530 535 540  
 Asn Tyr Gln Val Asp Glu Leu Ser Cys Glu Leu Cys Pro Leu Asp Gln  
 545 550 555 560

Arg Pro Asn Met Asn Arg Thr Gly Cys Gln Leu Ile Pro Ile Ile Lys  
 565 570 575  
 Leu Glu Trp His Ser Pro Trp Ala Val Val Pro Val Phe Val Ala Ile  
 580 585 590  
 Leu Gly Ile Ile Ala Thr Thr Phe Val Ile Val Thr Phe Val Arg Tyr  
 595 600 605  
 Asn Asp Thr Pro Ile Val Arg Ala Ser Gly Arg Glu Leu Ser Tyr Val  
 610 615 620  
 Leu Leu Thr Gly Ile Phe Leu Cys Tyr Ser Ile Thr Phe Leu Met Ile  
 625 630 635 640  
 Ala Ala Pro Asp Thr Ile Ile Cys Ser Phe Arg Arg Val Phe Leu Gly  
 645 650 655  
 Leu Gly Met Cys Phe Ser Tyr Ala Ala Leu Leu Thr Lys Thr Asn Arg  
 660 665 670  
 Ile His Arg Ile Phe Glu Gln Gly Lys Lys Ser Val Thr Ala Pro Lys  
 675 680 685  
 Phe Ile Ser Pro Ala Ser Gln Leu Val Ile Thr Phe Ser Leu Ile Ser  
 690 695 700  
 Val Gln Leu Leu Gly Val Phe Val Trp Phe Val Val Asp Pro Pro His  
 705 710 715 720  
 Ile Ile Ile Asp Tyr Gly Glu Gln Arg Thr Leu Asp Pro Glu Lys Ala  
 725 730 735  
 Arg Gly Val Leu Lys Cys Asp Ile Ser Asp Leu Ser Leu Ile Cys Ser  
 740 745 750  
 Leu Gly Tyr Ser Ile Leu Leu Met Val Thr Cys Thr Val Tyr Ala Ile  
 755 760 765  
 Lys Thr Arg Gly Val Pro Glu Thr Phe Asn Glu Ala Lys Pro Ile Gly  
 770 775 780  
 Phe Thr Met Tyr Thr Thr Cys Ile Ile Trp Leu Ala Phe Ile Pro Ile  
 785 790 795 800  
 Phe Phe Gly Thr Ala Gln Ser Ala Glu Lys Met Tyr Ile Gln Thr Thr  
 805 810 815  
 Thr Leu Thr Val Ser Met Ser Leu Ser Ala Ser Val Ser Leu Gly Met  
 820 825 830  
 Leu Tyr Met Pro Lys Val Tyr Ile Ile Ile Phe His Pro Glu Gln Asn  
 835 840 845  
 Thr Ile Glu Glu Val Arg Cys Ser Thr Ala Ala His Ala Phe Lys Val



850                      855                      860  
 Ala Ala Arg Ala Thr Leu Arg Arg Ser Asn Val Ser Arg Lys Arg Ser  
 865                      870                      875                      880  
 Ser Ser Leu Gly Gly Ser Thr Gly Ser Thr Pro Ser Ser Ser Ile Ser  
                     885                      890                      895  
 Ser Lys Ser Asn Ser Glu Asp Pro Phe Pro Gln Pro Glu Arg Gln Lys  
                     900                      905                      910  
 Gln Gln Gln Pro Leu Ala Leu Thr Gln Gln Glu Gln Gln Gln Gln Pro  
                     915                      920                      925  
 Leu Thr Leu Pro Gln Gln Gln Arg Ser Gln Gln Gln Pro Arg Cys Lys  
                     930                      935                      940  
 Gln Lys Val Ile Phe Gly Ser Gly Thr Val Thr Phe Ser Leu Ser Phe  
 945                      950                      955                      960  
 Asp Glu Pro Gln Lys Asn Ala Met Ala His Gly Asn Ser Thr His Gln  
                     965                      970                      975  
 Asn Ser Leu Glu Ala Gln Lys Ser Ser Asp Thr Leu Thr Arg His Gln  
                     980                      985                      990  
 Pro Leu Leu Pro Leu Gln Cys Gly Glu Thr Asp Leu Asp Leu Thr Val  
                     995                      1000                      1005  
 Gln Glu Thr Gly Leu Gln Gly Pro Val Gly Gly Asp Gln Arg Pro Glu  
                     1010                      1015                      1020  
 Val Glu Asp Pro Glu Glu Leu Ser Pro Ala Leu Val Val Ser Ser Ser  
 1025                      1030                      1035                      1040  
 Gln Ser Phe Val Ile Ser Gly Gly Gly Ser Thr Val Thr Glu Asn Val  
                     1045                      1050                      1055  
 Val Asn Ser

&lt;210&gt; 40

&lt;211&gt; 4257

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; Chimeric mGluR8//CaR\*G(qi5

&lt;400&gt; 40

atggtatgcg agggaaagcg atcagcctct tgcccttggt tcttcctctt gaccgccaag      60  
 ttctactgga tctcacaat gatgcaaaga actcacagcc aggagtatgc ccattccata      120  
 cgggtggatg gggacattat tttggggggt ctcttccttg tccacgcaaa gggagagaga      180

ggggtgcctt	gtggggagct	gaagaaggaa	aaggggattc	acagactgga	ggccatgctt	240
tatgcaattg	accagattaa	caaggaccct	gatctccttt	ccaacatcac	tctgggtgtc	300
cgcacctctg	acacgtgctc	tagggacacc	tatgcttttg	agcagtctct	aacattcgtg	360
caggcattaa	tagagaaaga	tgcttcggat	gtgaagtgtg	ctaattggaga	tccacccatt	420
ttcaccaagc	ccgacaagat	ttctggcgtc	ataggtgctg	cagcaagctc	cgtgtccatc	480
atggttgcta	acattttaag	actttttaag	atacctcaaa	tcagctatgc	atccacagcc	540
ccagagctaa	gtgataacac	caggtatgac	ttttctctc	gagtggttcc	gcctgactcc	600
taccaagccc	aagccatggt	ggacatcgtg	acagcactgg	gatggaatta	tgtttcgaca	660
ctggcttctg	aggggaacta	tggtagagc	ggtgtggagg	ccttcaccca	gatctcgagg	720
gagattggtg	gtgtttgcat	tgctcagtc	cagaaaaatcc	cacgtgaacc	aagacctgga	780
gaatttgaaa	aaattatcaa	acgcctgcta	gaaacaccta	atgctcgagc	agtgattatg	840
tttgccaatg	aggatgacat	caggaggata	ttggaagcag	caaaaaaact	aaaccaaaagt	900
gggcattttc	tctggattgg	ctcagatagt	tggggatcca	aaatagcacc	tgtctatcag	960
caagaggaga	ttgcagaagg	ggctgtgaca	attttgccca	aacgagcatc	aattgatgga	1020
tttgatcgat	actttagaag	ccgaactctt	gccaataatc	gaagaaatgt	gtggtttgca	1080
gaattctggg	aggagaattt	tggctgcaag	ttaggatcac	atgggaaaag	gaacagtcac	1140
ataaagaaat	gcacagggct	ggagcgaatt	gctcgggatt	catcttatga	acaggaagga	1200
aaggtccaat	ttgtaattga	tgctgtatat	tccatggctt	acgccttgca	caatatgcac	1260
aaagatctct	gccctggata	cattggcctt	tgtccacgaa	tgagtaccat	tgatgggaaa	1320
gagctacttg	gttatattcg	ggctgtaaat	tttaatggca	gtgctggcac	tcctgtcact	1380
tttaatgaaa	acggagatgc	tcctggacgt	tatgatatct	tccagtatca	aataaccaac	1440
aaaagcacag	agtacaaagt	catcgccac	tggaccaatc	agcttcatct	aaaagtggaa	1500
gacatgcagt	gggctcatag	agaacatact	cacccggcgt	ctgtctgcag	cctgccgtgt	1560
aagccagggg	agaggaagaa	aacggtgaaa	ggggtccctt	gctgctggca	ctgtgaacgc	1620
tgtgaagggt	acaactacca	ggtggatgag	ctgtcctgtg	aactttgccc	tctggatcag	1680
agaccaaca	tgaaccgcac	aggctgccag	cttatcccca	tcatacaaat	ggagtggcat	1740
tctccctggg	ctgtggtgcc	tgtgtttggt	gcaatatggg	gaatcatcgc	caccaccttt	1800
gtgatcgtga	cctttgtccg	ctataatgac	acacctatcg	tgagggcctc	aggacgcgaa	1860
cttagttacg	tgtccttaac	ggggattttt	cctctgtatt	caatcacgtt	tttaatgatt	1920
gcagcaccag	atacaatcat	atgctccttc	cgacgggtct	tcctaggact	tggcatgtgt	1980
ttcagctatg	cagcccttct	gacaaaaaca	aaccgtatcc	accgaatatt	tgagcagggg	2040
aagaaatctg	tcacagcgcc	caagttcatt	agtccagcat	ctcagctggt	gatcaccttc	2100
agcctcatct	cgtccagct	ccttgagtg	tttgtctggt	ttgttgtgga	tcaccccccac	2160
atcatcattg	actatggaga	gcagcggaca	ctagatccag	agaaggccag	gggagtgtct	2220
aagtgtgaca	tttctgatct	ctcactcatt	tgttcacttg	gatacagtat	cctcttgatg	2280
gtcacttgta	ctgtttatgc	cattaaaacg	agaggtgtcc	cagagacttt	caatgaagcc	2340
aaacctattg	gatttaccat	gtataaccac	tgcactatct	ggttagcttt	catccccatc	2400
ttttttggta	cagcccagtc	agcagaaaag	atgtacatcc	agacaacaac	acttactgtc	2460
tccatgagtt	taagtgtctc	agtatctctg	ggcatgctct	atatgcccaa	ggtttatatt	2520
ataatttttc	atccagaaca	gaataccatc	gaggaggtgc	gttgacgac	cgcagctcac	2580
gctttcaagg	tggctgcccg	ggccacgctg	cgccgcagca	acgtctcccg	caagcgggtcc	2640
agcagccttg	gaggetccac	gggatccacc	ccctcctcct	ccatcagcag	caagagcaac	2700
agcgaagacc	cattcccaca	gcccagagag	cagaagcagc	agcagccgct	ggccctaacc	2760
cagcaagagc	agcagcagca	gcccctgacc	ctcccacagc	agcaacgac	tcagcagcag	2820
cccagatgca	agcagaaggt	catctttggc	agcggcacgg	tcaccttctc	actgagcttt	2880
gatgagcctc	agaagaacgc	catggcccac	gggaattcta	cgcaccagaa	ctccctggag	2940
gcccagaaaa	gcagcgatac	gctgacccga	caccagccat	tactcccgtc	gcagtgcggg	3000
gaaacggact	tagatctgac	cgtccaggaa	acaggtctgc	aaggacctgt	gggtggagac	3060
cagcggccag	aggtggagga	ccctgaagag	ttgtccccag	cacttgtagt	gtccagttca	3120
cagagctttg	tcacatagtg	tggaggcagc	actgttacag	aaaacgtagt	gaattcaatg	3180
actctggagt	ccatcatggc	gtgctgctg	agcagggagg	ccaaggaagc	ccggcggatc	3240
aacgacgaga	tcagcgggca	gctccgcagg	gacaagcggg	acgcccggcg	ggagctcaag	3300
ctgctgctgc	tcgggacagg	agagagtggc	aagagtacgt	ttatcaagca	gatgagaatc	3360
atccatgggt	caggatactc	tgatgaagat	aaaaggggct	tcaccaagct	ggtgtatcag	3420
aacatcttca	cggccatgca	ggccatgatc	agagccatgg	acacactcaa	gatcccatac	3480
aagtatgagc	acaataaggc	tcatgcacaa	ttagtctcag	aagttgatgt	ggagaagggtg	3540
tctgcttttg	agaatccata	tgtagatgca	ataaagagtt	tatggaatga	tcctggaatc	3600

```

caggaatgct atgatagacg acgagaatat caattatctg actctaccaa atactatctt 3660
aatgacttgg accgcgtagc tgaccctgcc tacctgccta cgcaacaaga tgtgcttaga 3720
gttcgagtcc ccaccacagg gatcatcgaa tacccttttg acttacaaag tgtcattttc 3780
agaatggtcg atgtaggggg ccaaagggtca gagagaagaa aatggatata ctgctttgaa 3840
aatgtcacct ctatcatgtt tctagtagcg cttagtgaat atgatcaagt tctcgtggag 3900
tcagacaatg agaaccgaat ggaggaaagc aaggctctct ttagaacaat tatcacatac 3960
ccctgggttc agaactcttc ggttattctg ttcttaaaca agaaagatct tctagaggag 4020
aaaatcatgt attcccatct agtcgactac ttcccagaat atgatggacc ccagagagat 4080
gcccaggcag cccgagaatt cattctgaag atgttcgtgg acctgaacct agacagtgac 4140
aaaattatct actcccactt cactgtcgcc acagacaccg agaatatccg ctttgtcttt 4200
gctgccgtca aggacaccat cctccagttg aacctgaagg actgcggtct gttcttaa 4257

```

<210> 41

<211> 1418

<212> PRT

<213> Artificial Sequence

<220>

<223> Chimeric mGluR8//CaR\*G(qi5

<400> 41

```

Met Val Cys Glu Gly Lys Arg Ser Ala Ser Cys Pro Cys Phe Phe Leu
 1              5              10              15

Leu Thr Ala Lys Phe Tyr Trp Ile Leu Thr Met Met Gln Arg Thr His
      20              25              30

Ser Gln Glu Tyr Ala His Ser Ile Arg Val Asp Gly Asp Ile Ile Leu
      35              40              45

Gly Gly Leu Phe Pro Val His Ala Lys Gly Glu Arg Gly Val Pro Cys
 50              55              60

Gly Glu Leu Lys Lys Glu Lys Gly Ile His Arg Leu Glu Ala Met Leu
 65              70              75              80

Tyr Ala Ile Asp Gln Ile Asn Lys Asp Pro Asp Leu Leu Ser Asn Ile
      85              90              95

Thr Leu Gly Val Arg Ile Leu Asp Thr Cys Ser Arg Asp Thr Tyr Ala
      100              105              110

Leu Glu Gln Ser Leu Thr Phe Val Gln Ala Leu Ile Glu Lys Asp Ala
      115              120              125

Ser Asp Val Lys Cys Ala Asn Gly Asp Pro Pro Ile Phe Thr Lys Pro
      130              135              140

Asp Lys Ile Ser Gly Val Ile Gly Ala Ala Ala Ser Ser Val Ser Ile
      145              150              155              160

Met Val Ala Asn Ile Leu Arg Leu Phe Lys Ile Pro Gln Ile Ser Tyr
      165              170              175

Ala Ser Thr Ala Pro Glu Leu Ser Asp Asn Thr Arg Tyr Asp Phe Phe
      180              185              190

```

Ser Arg Val Val Pro Pro Asp Ser Tyr Gln Ala Gln Ala Met Val Asp  
 195 200 205  
 Ile Val Thr Ala Leu Gly Trp Asn Tyr Val Ser Thr Leu Ala Ser Glu  
 210 215 220  
 Gly Asn Tyr Gly Glu Ser Gly Val Glu Ala Phe Thr Gln Ile Ser Arg  
 225 230 235 240  
 Glu Ile Gly Gly Val Cys Ile Ala Gln Ser Gln Lys Ile Pro Arg Glu  
 245 250 255  
 Pro Arg Pro Gly Glu Phe Glu Lys Ile Ile Lys Arg Leu Leu Glu Thr  
 260 265 270  
 Pro Asn Ala Arg Ala Val Ile Met Phe Ala Asn Glu Asp Asp Ile Arg  
 275 280 285  
 Arg Ile Leu Glu Ala Ala Lys Lys Leu Asn Gln Ser Gly His Phe Leu  
 290 295 300  
 Trp Ile Gly Ser Asp Ser Trp Gly Ser Lys Ile Ala Pro Val Tyr Gln  
 305 310 315 320  
 Gln Glu Glu Ile Ala Glu Gly Ala Val Thr Ile Leu Pro Lys Arg Ala  
 325 330 335  
 Ser Ile Asp Gly Phe Asp Arg Tyr Phe Arg Ser Arg Thr Leu Ala Asn  
 340 345 350  
 Asn Arg Arg Asn Val Trp Phe Ala Glu Phe Trp Glu Glu Asn Phe Gly  
 355 360 365  
 Cys Lys Leu Gly Ser His Gly Lys Arg Asn Ser His Ile Lys Lys Cys  
 370 375 380  
 Thr Gly Leu Glu Arg Ile Ala Arg Asp Ser Ser Tyr Glu Gln Glu Gly  
 385 390 395 400  
 Lys Val Gln Phe Val Ile Asp Ala Val Tyr Ser Met Ala Tyr Ala Leu  
 405 410 415  
 His Asn Met His Lys Asp Leu Cys Pro Gly Tyr Ile Gly Leu Cys Pro  
 420 425 430  
 Arg Met Ser Thr Ile Asp Gly Lys Glu Leu Leu Gly Tyr Ile Arg Ala  
 435 440 445  
 Val Asn Phe Asn Gly Ser Ala Gly Thr Pro Val Thr Phe Asn Glu Asn  
 450 455 460  
 Gly Asp Ala Pro Gly Arg Tyr Asp Ile Phe Gln Tyr Gln Ile Thr Asn  
 465 470 475 480

Lys Ser Thr Glu Tyr Lys Val Ile Gly His Trp Thr Asn Gln Leu His  
 485 490 495  
 Leu Lys Val Glu Asp Met Gln Trp Ala His Arg Glu His Thr His Pro  
 500 505 510  
 Ala Ser Val Cys Ser Leu Pro Cys Lys Pro Gly Glu Arg Lys Lys Thr  
 515 520 525  
 Val Lys Gly Val Pro Cys Cys Trp His Cys Glu Arg Cys Glu Gly Tyr  
 530 535 540  
 Asn Tyr Gln Val Asp Glu Leu Ser Cys Glu Leu Cys Pro Leu Asp Gln  
 545 550 555 560  
 Arg Pro Asn Met Asn Arg Thr Gly Cys Gln Leu Ile Pro Ile Ile Lys  
 565 570 575  
 Leu Glu Trp His Ser Pro Trp Ala Val Val Pro Val Phe Val Ala Ile  
 580 585 590  
 Leu Gly Ile Ile Ala Thr Thr Phe Val Ile Val Thr Phe Val Arg Tyr  
 595 600 605  
 Asn Asp Thr Pro Ile Val Arg Ala Ser Gly Arg Glu Leu Ser Tyr Val  
 610 615 620  
 Leu Leu Thr Gly Ile Phe Leu Cys Tyr Ser Ile Thr Phe Leu Met Ile  
 625 630 635 640  
 Ala Ala Pro Asp Thr Ile Ile Cys Ser Phe Arg Arg Val Phe Leu Gly  
 645 650 655  
 Leu Gly Met Cys Phe Ser Tyr Ala Ala Leu Leu Thr Lys Thr Asn Arg  
 660 665 670  
 Ile His Arg Ile Phe Glu Gln Gly Lys Lys Ser Val Thr Ala Pro Lys  
 675 680 685  
 Phe Ile Ser Pro Ala Ser Gln Leu Val Ile Thr Phe Ser Leu Ile Ser  
 690 695 700  
 Val Gln Leu Leu Gly Val Phe Val Trp Phe Val Val Asp Pro Pro His  
 705 710 715 720  
 Ile Ile Ile Asp Tyr Gly Glu Gln Arg Thr Leu Asp Pro Glu Lys Ala  
 725 730 735  
 Arg Gly Val Leu Lys Cys Asp Ile Ser Asp Leu Ser Leu Ile Cys Ser  
 740 745 750  
 Leu Gly Tyr Ser Ile Leu Leu Met Val Thr Cys Thr Val Tyr Ala Ile  
 755 760 765  
 Lys Thr Arg Gly Val Pro Glu Thr Phe Asn Glu Ala Lys Pro Ile Gly  
 770 775 780

Phe Thr Met Tyr Thr Thr Cys Ile Ile Trp Leu Ala Phe Ile Pro Ile  
785 790 795 800

Phe Phe Gly Thr Ala Gln Ser Ala Glu Lys Met Tyr Ile Gln Thr Thr  
805 810 815

Thr Leu Thr Val Ser Met Ser Leu Ser Ala Ser Val Ser Leu Gly Met  
820 825 830

Leu Tyr Met Pro Lys Val Tyr Ile Ile Ile Phe His Pro Glu Gln Asn  
835 840 845

Thr Ile Glu Glu Val Arg Cys Ser Thr Ala Ala His Ala Phe Lys Val  
850 855 860

Ala Ala Arg Ala Thr Leu Arg Arg Ser Asn Val Ser Arg Lys Arg Ser  
865 870 875 880

Ser Ser Leu Gly Gly Ser Thr Gly Ser Thr Pro Ser Ser Ser Ile Ser  
885 890 895

Ser Lys Ser Asn Ser Glu Asp Pro Phe Pro Gln Pro Glu Arg Gln Lys  
900 905 910

Gln Gln Gln Pro Leu Ala Leu Thr Gln Gln Glu Gln Gln Gln Pro  
915 920 925

Leu Thr Leu Pro Gln Gln Gln Arg Ser Gln Gln Gln Pro Arg Cys Lys  
930 935 940

Gln Lys Val Ile Phe Gly Ser Gly Thr Val Thr Phe Ser Leu Ser Phe  
945 950 955 960

Asp Glu Pro Gln Lys Asn Ala Met Ala His Gly Asn Ser Thr His Gln  
965 970 975

Asn Ser Leu Glu Ala Gln Lys Ser Ser Asp Thr Leu Thr Arg His Gln  
980 985 990

Pro Leu Leu Pro Leu Gln Cys Gly Glu Thr Asp Leu Asp Leu Thr Val  
995 1000 1005

Gln Glu Thr Gly Leu Gln Gly Pro Val Gly Gly Asp Gln Arg Pro Glu  
1010 1015 1020

Val Glu Asp Pro Glu Glu Leu Ser Pro Ala Leu Val Val Ser Ser Ser  
1025 1030 1035 1040

Gln Ser Phe Val Ile Ser Gly Gly Gly Ser Thr Val Thr Glu Asn Val  
1045 1050 1055

Val Asn Ser Met Thr Leu Glu Ser Ile Met Ala Cys Cys Leu Ser Glu  
1060 1065 1070

Glu Ala Lys Glu Ala Arg Arg Ile Asn Asp Glu Ile Glu Arg Gln Leu

1075	1080	1085
Arg Arg Asp Lys Arg Asp 1090	Ala Arg Arg Glu Leu 1095	Lys Leu Leu Leu Leu 1100
Gly Thr Gly Glu Ser 1105	Gly Lys Ser Thr Phe 1110	Ile Lys Gln Met Arg Ile 1115 1120
Ile His Gly Ser Gly Tyr Ser Asp 1125	Glu Asp Lys Arg Gly Phe Thr 1130 1135	
Leu Val Tyr Gln Asn Ile Phe Thr 1140	Ala Met Gln Ala Met 1145 1150	Ile Arg Ala
Met Asp Thr Leu Lys Ile Pro Tyr Lys Tyr Glu His Asn Lys Ala His 1155 1160 1165		
Ala Gln Leu Val Arg Glu Val Asp Val Glu Lys Val Ser Ala Phe Glu 1170 1175 1180		
Asn Pro Tyr Val Asp Ala Ile Lys Ser Leu Trp Asn Asp Pro Gly Ile 1185 1190 1195 1200		
Gln Glu Cys Tyr Asp Arg Arg Arg Glu Tyr Gln Leu Ser Asp Ser Thr 1205 1210 1215		
Lys Tyr Tyr Leu Asn Asp Leu Asp Arg Val Ala Asp Pro Ala Tyr Leu 1220 1225 1230		
Pro Thr Gln Gln Asp Val Leu Arg Val Arg Val Pro Thr Thr Gly Ile 1235 1240 1245		
Ile Glu Tyr Pro Phe Asp Leu Gln Ser Val Ile Phe Arg Met Val Asp 1250 1255 1260		
Val Gly Gly Gln Arg Ser Glu Arg Arg Lys Trp Ile His Cys Phe Glu 1265 1270 1275 1280		
Asn Val Thr Ser Ile Met Phe Leu Val Ala Leu Ser Glu Tyr Asp Gln 1285 1290 1295		
Val Leu Val Glu Ser Asp Asn Glu Asn Arg Met Glu Glu Ser Lys Ala 1300 1305 1310		
Leu Phe Arg Thr Ile Ile Thr Tyr Pro Trp Phe Gln Asn Ser Ser Val 1315 1320 1325		
Ile Leu Phe Leu Asn Lys Lys Asp Leu Leu Glu Glu Lys Ile Met Tyr 1330 1335 1340		
Ser His Leu Val Asp Tyr Phe Pro Glu Tyr Asp Gly Pro Gln Arg Asp 1345 1350 1355 1360		
Ala Gln Ala Ala Arg Glu Phe Ile Leu Lys Met Phe Val Asp Leu Asn 1365 1370 1375		

```
<210> 42
<211> 3909
<212> DNA
<213> Artificial Sequence

<220>
<223> Chimeric GABA-BR2*Gqo5

<400> 42
```

atggcttccc	cgcggagctc	cgggcagccc	gggcgcgcgc	cgccgcgcgc	accgcgcgcc	60
cgcgcctgc	tactgtact	gctgctgccg	ctgctgctgc	ctctggcgcc	cggggcctgg	120
ggctgggcgc	ggggcgcccc	cgggcgcgcg	cccagcagcc	cgcgcctctc	catcatgggc	180
ctcatgccgc	taccaagga	ggtggccaag	ggcagcatcg	ggcgcggtgt	gctccccgcc	240
gtggaactgg	ccatcgagca	gatccgcaac	gagtcactcc	tgcgccctta	cttcctcgac	300
ctgcggtctc	atgacacgga	gtgcgacaac	gcaaaaggtt	tgaagcctt	ctacgatgca	360
ataaaaatacg	ggccgaacca	cttgatggtg	tttggaggcg	tctgtccatc	cgtcacatcc	420
atcattgcag	agtcctccca	aggctggaat	ctggtgcagc	tttcttttgc	tgcaaccacg	480
cctgttctag	ccgataagaa	aaaataccct	tatttctttc	ggaccgtccc	atcagacaat	540
gcgtgtaatc	cagccattct	gaagttgctc	aagcactacc	agtggaagcg	cgtgggcacg	600
ctgacgcaag	acgttcagag	gttctctgag	gtgcggaatg	acctgactgg	agttctgtat	660
ggcgaggaca	ttgagatttc	agacaccgag	agcttctcca	acgatccctg	taccagtgtc	720
aaaaagctga	aggggaatga	tgtgcggatc	atcettggcc	agtttgacca	gaatatggca	780
gcaaaagtgt	tctgtttgtc	atacgaggag	aacatglatg	gtagtaaata	tcagtggatc	840
attccgggct	ggtacgagcc	ttcttgggtg	gagcaggtgc	acacggaagc	caactcatcc	900
cgctgccttc	ggaagaatct	gcttgtgtgc	atggaggctc	acattggcgt	ggatttcgag	960
ccctgagct	ccaagcagat	caagaccatc	tcaggaaaga	ctccacagca	gstatgagaga	1020
gagtacaaca	acaagcggtc	aggcgtgggg	cccagcaagt	tccacgggta	cgctacgat	1080
ggcatctggg	tcatcgccaa	gacactgcag	agggccatgg	agacactgca	tgccagcagc	1140
cggcaccagc	ggatccagga	cttcaactac	acggaccaca	cgtggggcag	gatcatcctc	1200
aatgccatga	acgagacca	cttcttcggg	gtcacgggtc	aagttgtatt	ccggaatggg	1260
gagagaatgg	ggaccattaa	atttactcaa	tttcaagaca	gcagggaggt	gaaggtggga	1320
gagtacaacg	ctgtggccga	cacactggag	atcatcaatg	acaccatcag	gttccaagga	1380
tccgaaccac	caaaagacaa	gaccatcatc	ctggagcagc	tgcggaagat	ctccctacct	1440
ctctacagca	tctctctctg	cctcaccatc	ctcgggatga	tcatggccag	tgtttttctc	1500
ttcttcaaca	tcaagaaccg	gaatcagaag	ctcataaaga	tgtcaggtcc	atacatgaac	1560
aaccttatca	tctttggagg	gatgctctcc	tatgcttcca	tatttctctt	tggccttgat	1620
ggatcctttg	tctctgaaaa	gacctttgaa	acacttttga	ccgtcaggac	ctggattctc	1680
accgtgggct	acacgaccgc	ttttggggcc	atgtttgcaa	agacctggag	agtcacagcc	1740
atcttcaaaa	atgtgaaaat	gaagaagaag	atcatcaagg	accagaaact	gcttgtgatc	1800
gtggggggca	tgtgtctgat	cgacctgtgt	atcctgatct	gctggcaggc	tgtggacccc	1860
ctgcgaagga	cagtggagaa	gtacagcatg	gagccggacc	cagcaggacg	ggatatctcc	1920
atccgccttc	tcttgaggca	ctgtgagaac	accatataga	ccaattggct	tggcatcgtc	1980
tatgcctaca	agggacttct	catgttgttc	ggttgtttct	tagcttggga	gaccgcgaac	2040
gtcagcatcc	ccgcactcaa	cgacagcaag	tacatcggga	tgagtgtcta	caacgtgggg	2100



```

atcatgtgca tcatcggggc cgctgtctcc ttctgaccc gggaccagcc caatgtgcag 2160
ttctgcatcg tggctctggt catcatcttc tgcagcacca tcacctctg cctggtattc 2220
gtgccgaagc tcatcaccct gagaacaaac ccagatgcag caacgcagaa caggcgattc 2280
cagttcactc agaatcagaa gaaagaagat tctaaaacgt ccacctcggg caccagtgtg 2340
aaccaagcca gcacatcccg cctggagggc ctacagtcag aaaaccatcg cctgcgaatg 2400
aagatcacag agctggataa agacttgga gaggtcacca tgcagctgca ggacacacca 2460
gaaaagacca cctacattaa acagaaccac taccaagagc tcaatgacat cctcaacctg 2520
ggaaacttca ctgagagcac agatggagga aaggccattt taaaaaatca cctcgatcaa 2580
aatccccagc tacagtggaa cacaacagag cctctctgaa catgcaaaga tcctatagaa 2640
gatataaact ctccagaaca catccagcgt cggctgtccc tccagctccc catcctccac 2700
cacgcctacc tcccattccat cggaggcgtg gacgccagct gtgtcagccc ctgcgtcagc 2760
cccaccgcca gccccgcca cagacatgtg ccacctctct tccgagtcac ggtctcgggc 2820
ctggcggcgc ccatgactct ggagtccatc atggcgtgct gcctgagcga ggaggccaag 2880
gaagcccggc ggatcaacga cgagatcgag cggcagctcc gcagggacaa gcgggacgcc 2940
cgccgggagc tcaagctgct gctgctcggg acaggagaga gtggcaagag tacgtttatc 3000
aagcagatga gaatcatcca tgggtcagga tactctgatg aagataaaaag gggcttcacc 3060
aagctggtgt atcagaacat cttcacggcc atgcaggcca tgatcagagc catggacaca 3120
ctcaagatcc catacaagta tgagcacaat aaggctcatg cacaattagt tgcagaagtt 3180
gatgtggaga aggtgtctgc ttttgagaat ccataatgtag atgcaataaa gagtttatgg 3240
aatgatcctg gaatccagga atgctatgat agacgacgag aatatcaatt atctgactct 3300
accaaatact atcttaatga cttggaccgc gtatgtgacc ctgcctacct gcctacgcaa 3360
caagatgtgc ttagagtctg agtccccacc acagggatca tgaatacccc ctttgactta 3420
caaagtgtca ttttcagaat ggtcgatgta gggggccaaa ggtcagagag aagaaaatgg 3480
atacactgct ttgaaaatgt cacctctatc atgtttctag tagcgttag tgaatatgat 3540
caagttctgg tggagtcaga caatgagaac cgaatggagg aaagcaaggc tctctttaga 3600
acaattatca catacccctg gttccagaac tcctcggtta ttctgttctt aaacaagaaa 3660
gatcttctag aggagaaaaat catgtattcc catctagtcg actacttccc agaatatgat 3720
ggaccccaga gagatgcccc ggcagcccga gaattcattc tgaagatggt cgtggacctg 3780
aaccagaca gtgacaaaaa taactactcc cacttcacgt gcgccacaga caccgagaat 3840
atccgctttg tctttgctgc cgtcaaggac accatcctcc agttgaacct gaagggtgcg 3900
ggtctgtac 3909

```

<210> 43

<211> 1303

<212> PRT

<213> Artificial Sequence

<220>

<223> Chimeric GABA-BR2\*Gqo5

<400> 43

```

Met Ala Ser Pro Arg Ser Ser Gly Gln Pro Gly Pro Pro Pro Pro Pro
1           5           10          15

Pro Pro Pro Pro Ala Arg Leu Leu Leu Leu Leu Leu Leu Pro Leu Leu
20          25          30

Leu Pro Leu Ala Pro Gly Ala Trp Gly Trp Ala Arg Gly Ala Pro Arg
35          40          45

Pro Pro Pro Ser Ser Pro Pro Leu Ser Ile Met Gly Leu Met Pro Leu
50          55          60

```

Thr Lys Glu Val Ala Lys Gly Ser Ile Gly Arg Gly Val Leu Pro Ala  
 65 70 75 80  
 Val Glu Leu Ala Ile Glu Gln Ile Arg Asn Glu Ser Leu Leu Arg Pro  
 85 90 95  
 Tyr Phe Leu Asp Leu Arg Leu Tyr Asp Thr Glu Cys Asp Asn Ala Lys  
 100 105 110  
 Gly Leu Lys Ala Phe Tyr Asp Ala Ile Lys Tyr Gly Pro Asn His Leu  
 115 120 125  
 Met Val Phe Gly Gly Val Cys Pro Ser Val Thr Ser Ile Ile Ala Glu  
 130 135 140  
 Ser Leu Gln Gly Trp Asn Leu Val Gln Leu Ser Phe Ala Ala Thr Thr  
 145 150 155 160  
 Pro Val Leu Ala Asp Lys Lys Lys Tyr Pro Tyr Phe Phe Arg Thr Val  
 165 170 175  
 Pro Ser Asp Asn Ala Val Asn Pro Ala Ile Leu Lys Leu Leu Lys His  
 180 185 190  
 Tyr Gln Trp Lys Arg Val Gly Thr Leu Thr Gln Asp Val Gln Arg Phe  
 195 200 205  
 Ser Glu Val Arg Asn Asp Leu Thr Gly Val Leu Tyr Gly Glu Asp Ile  
 210 215 220  
 Glu Ile Ser Asp Thr Glu Ser Phe Ser Asn Asp Pro Cys Thr Ser Val  
 225 230 235 240  
 Lys Lys Leu Lys Gly Asn Asp Val Arg Ile Ile Leu Gly Gln Phe Asp  
 245 250 255  
 Gln Asn Met Ala Ala Lys Val Phe Cys Cys Ala Tyr Glu Glu Asn Met  
 260 265 270  
 Tyr Gly Ser Lys Tyr Gln Trp Ile Ile Pro Gly Trp Tyr Glu Pro Ser  
 275 280 285  
 Trp Trp Glu Gln Val His Thr Glu Ala Asn Ser Ser Arg Cys Leu Arg  
 290 295 300  
 Lys Asn Leu Leu Ala Ala Met Glu Gly Tyr Ile Gly Val Asp Phe Glu  
 305 310 315 320  
 Pro Leu Ser Ser Lys Gln Ile Lys Thr Ile Ser Gly Lys Thr Pro Gln  
 325 330 335  
 Gln Tyr Glu Arg Glu Tyr Asn Asn Lys Arg Ser Gly Val Gly Pro Ser  
 340 345 350  
 Lys Phe His Gly Tyr Ala Tyr Asp Gly Ile Trp Val Ile Ala Lys Thr  
 355 360 365

Leu Gln Arg Ala Met Glu Thr Leu His Ala Ser Ser Arg His Gln Arg  
 370 375 380  
 Ile Gln Asp Phe Asn Tyr Thr Asp His Thr Leu Gly Arg Ile Ile Leu  
 385 390 395 400  
 Asn Ala Met Asn Glu Thr Asn Phe Phe Gly Val Thr Gly Gln Val Val  
 405 410 415  
 Phe Arg Asn Gly Glu Arg Met Gly Thr Ile Lys Phe Thr Gln Phe Gln  
 420 425 430  
 Asp Ser Arg Glu Val Lys Val Gly Glu Tyr Asn Ala Val Ala Asp Thr  
 435 440 445  
 Leu Glu Ile Ile Asn Asp Thr Ile Arg Phe Gln Gly Ser Glu Pro Pro  
 450 455 460  
 Lys Asp Lys Thr Ile Ile Leu Glu Gln Leu Arg Lys Ile Ser Leu Pro  
 465 470 475 480  
 Leu Tyr Ser Ile Leu Ser Ala Leu Thr Ile Leu Gly Met Ile Met Ala  
 485 490 495  
 Ser Ala Phe Leu Phe Phe Asn Ile Lys Asn Arg Asn Gln Lys Leu Ile  
 500 505 510  
 Lys Met Ser Ser Pro Tyr Met Asn Asn Leu Ile Ile Leu Gly Gly Met  
 515 520 525  
 Leu Ser Tyr Ala Ser Ile Phe Leu Phe Gly Leu Asp Gly Ser Phe Val  
 530 535 540  
 Ser Glu Lys Thr Phe Glu Thr Leu Cys Thr Val Arg Thr Trp Ile Leu  
 545 550 555 560  
 Thr Val Gly Tyr Thr Thr Ala Phe Gly Ala Met Phe Ala Lys Thr Trp  
 565 570 575  
 Arg Val His Ala Ile Phe Lys Asn Val Lys Met Lys Lys Lys Ile Ile  
 580 585 590  
 Lys Asp Gln Lys Leu Leu Val Ile Val Gly Gly Met Leu Leu Ile Asp  
 595 600 605  
 Leu Cys Ile Leu Ile Cys Trp Gln Ala Val Asp Pro Leu Arg Arg Thr  
 610 615 620  
 Val Glu Lys Tyr Ser Met Glu Pro Asp Pro Ala Gly Arg Asp Ile Ser  
 625 630 635 640  
 Ile Arg Pro Leu Leu Glu His Cys Glu Asn Thr His Met Thr Ile Trp  
 645 650 655  
 Leu Gly Ile Val Tyr Ala Tyr Lys Gly Leu Leu Met Leu Phe Gly Cys

660					665					670						
Phe	Leu	Ala	Trp	Glu	Thr	Arg	Asn	Val	Ser	Ile	Pro	Ala	Leu	Asn	Asp	
675					680					685						
Ser	Lys	Tyr	Ile	Gly	Met	Ser	Val	Tyr	Asn	Val	Gly	Ile	Met	Cys	Ile	
690					695					700						
Ile	Gly	Ala	Ala	Val	Ser	Phe	Leu	Thr	Arg	Asp	Gln	Pro	Asn	Val	Gln	
705					710					715					720	
Phe	Cys	Ile	Val	Ala	Leu	Val	Ile	Ile	Phe	Cys	Ser	Thr	Ile	Thr	Leu	
725					730					735						
Cys	Leu	Val	Phe	Val	Pro	Lys	Leu	Ile	Thr	Leu	Arg	Thr	Asn	Pro	Asp	
740					745					750						
Ala	Ala	Thr	Gln	Asn	Arg	Arg	Phe	Gln	Phe	Thr	Gln	Asn	Gln	Lys	Lys	
755					760					765						
Glu	Asp	Ser	Lys	Thr	Ser	Thr	Ser	Val	Thr	Ser	Val	Asn	Gln	Ala	Ser	
770					775					780						
Thr	Ser	Arg	Leu	Glu	Gly	Leu	Gln	Ser	Glu	Asn	His	Arg	Leu	Arg	Met	
785					790					795					800	
Lys	Ile	Thr	Glu	Leu	Asp	Lys	Asp	Leu	Glu	Glu	Val	Thr	Met	Gln	Leu	
805					810					815						
Gln	Asp	Thr	Pro	Glu	Lys	Thr	Thr	Tyr	Ile	Lys	Gln	Asn	His	Tyr	Gln	
820					825					830						
Glu	Leu	Asn	Asp	Ile	Leu	Asn	Leu	Gly	Asn	Phe	Thr	Glu	Ser	Thr	Asp	
835					840					845						
Gly	Gly	Lys	Ala	Ile	Leu	Lys	Asn	His	Leu	Asp	Gln	Asn	Pro	Gln	Leu	
850					855					860						
Gln	Trp	Asn	Thr	Thr	Glu	Pro	Ser	Arg	Thr	Cys	Lys	Asp	Pro	Ile	Glu	
865					870					875					880	
Asp	Ile	Asn	Ser	Pro	Glu	His	Ile	Gln	Arg	Arg	Leu	Ser	Leu	Gln	Leu	
885					890					895						
Pro	Ile	Leu	His	His	Ala	Tyr	Leu	Pro	Ser	Ile	Gly	Gly	Val	Asp	Ala	
900					905					910						
Ser	Cys	Val	Ser	Pro	Cys	Val	Ser	Pro	Thr	Ala	Ser	Pro	Arg	His	Arg	
915					920					925						
His	Val	Pro	Pro	Ser	Phe	Arg	Val	Met	Val	Ser	Gly	Leu	Ala	Ala	Ala	
930					935					940						
Met	Thr	Leu	Glu	Ser	Ile	Met	Ala	Cys	Cys	Leu	Ser	Glu	Glu	Ala	Lys	
945					950					955					960	

Glu Ala Arg Arg Ile Asn Asp Glu Ile Glu Arg Gln Leu Arg Arg Asp  
 965 970 975  
 Lys Arg Asp Ala Arg Arg Glu Leu Lys Leu Leu Leu Leu Gly Thr Gly  
 980 985 990  
 Glu Ser Gly Lys Ser Thr Phe Ile Lys Gln Met Arg Ile Ile His Gly  
 995 1000 1005  
 Ser Gly Tyr Ser Asp Glu Asp Lys Arg Gly Phe Thr Lys Leu Val Tyr  
 1010 1015 1020  
 Gln Asn Ile Phe Thr Ala Met Gln Ala Met Ile Arg Ala Met Asp Thr  
 1025 1030 1035 1040  
 Leu Lys Ile Pro Tyr Lys Tyr Glu His Asn Lys Ala His Ala Gln Leu  
 1045 1050 1055  
 Val Arg Glu Val Asp Val Glu Lys Val Ser Ala Phe Glu Asn Pro Tyr  
 1060 1065 1070  
 Val Asp Ala Ile Lys Ser Leu Trp Asn Asp Pro Gly Ile Gln Glu Cys  
 1075 1080 1085  
 Tyr Asp Arg Arg Arg Glu Tyr Gln Leu Ser Asp Ser Thr Lys Tyr Tyr  
 1090 1095 1100  
 Leu Asn Asp Leu Asp Arg Val Ala Asp Pro Ala Tyr Leu Pro Thr Gln  
 1105 1110 1115 1120  
 Gln Asp Val Leu Arg Val Arg Val Pro Thr Thr Gly Ile Ile Glu Tyr  
 1125 1130 1135  
 Pro Phe Asp Leu Gln Ser Val Ile Phe Arg Met Val Asp Val Gly Gly  
 1140 1145 1150  
 Gln Arg Ser Glu Arg Arg Lys Trp Ile His Cys Phe Glu Asn Val Thr  
 1155 1160 1165  
 Ser Ile Met Phe Leu Val Ala Leu Ser Glu Tyr Asp Gln Val Leu Val  
 1170 1175 1180  
 Glu Ser Asp Asn Glu Asn Arg Met Glu Glu Ser Lys Ala Leu Phe Arg  
 1185 1190 1195 1200  
 Thr Ile Ile Thr Tyr Pro Trp Phe Gln Asn Ser Ser Val Ile Leu Phe  
 1205 1210 1215  
 Leu Asn Lys Lys Asp Leu Leu Glu Glu Lys Ile Met Tyr Ser His Leu  
 1220 1225 1230  
 Val Asp Tyr Phe Pro Glu Tyr Asp Gly Pro Gln Arg Asp Ala Gln Ala  
 1235 1240 1245  
 Ala Arg Glu Phe Ile Leu Lys Met Phe Val Asp Leu Asn Pro Asp Ser  
 1250 1255 1260

Asp Lys Ile Asn Tyr Ser His Phe Thr Cys Ala Thr Asp Thr Glu Asn  
 1265 1270 1275 1280

Ile Arg Phe Val Phe Ala Ala Val Lys Asp Thr Ile Leu Gln Leu Asn  
 1285 1290 1295

Leu Lys Gly Cys Gly Leu Tyr  
 1300

<210> 44

<211> 3969

<212> DNA

<213> Artificial Sequence

<220>

<223> Chimeric GABA-BR1a\*Gqo5

<400> 44

atgttgc	tgctgctact	ggcgccactc	ttcctccgcc	ccccgggcgc	ggcgggggcg	60
cagaccccc	acgccacctc	agaaggttgc	cagatcatat	acccgccctg	ggaagggggc	120
atcaggtacc	ggggcctgac	tcgggaccag	gtgaaggcta	tcaacttctt	gccagtggac	180
tatgagattg	agtatgtgtg	ccggggggag	cgcgaggtgg	tggggcccaa	ggcccgcaag	240
tgcttgccca	acggctcctg	gacagatatg	gacacaccca	gccgctgtgt	ccgaatctgc	300
tccaagtctt	atttgaccct	ggaaaaatgg	aaggttttcc	tgacgggtgg	ggacctccca	360
gctctggacg	gagcccgggt	ggatttcggg	tgtgaccccg	acttccatct	gggtgggcagc	420
tcccggagca	tctgtagtca	gggccagtg	agcaccacca	agccccactg	ccaggtgaat	480
cgaacgccac	actcagaacg	gcgcgcagtg	tacatcgggg	cactgtttcc	catgagcggg	540
ggctggccag	ggggccaggc	ctgccagccc	gcggtggaga	tggcgctgga	ggacgtgaat	600
agccgcaggg	acatcctgcc	ggactatgag	ctcaagctca	tccaccacga	cagcaagtgt	660
gatccaggcc	aagccacca	gtacctatat	gagctgctct	acaacgaccc	tatcaagatc	720
atccttatgc	ctggctgcag	ctctgtctcc	acgctggtgg	ctgaggctgc	taggatgtgg	780
aacctcattg	tgcttttcta	tggtctccagc	tcaccagccc	tgtcaaaccg	gcagcgtttc	840
cccactttct	tccgaacgca	cccacagcc	acactccaca	accctacccg	cgtgaaactc	900
tttgaaaagt	ggggctggaa	gaagattgct	accatccagc	agaccactga	ggtcttccact	960
togactctgg	acgacctgga	ggaacgagtg	aaggaggctg	gaattgagat	tactttccgc	1020
cagagtttct	tctcagatcc	agctgtgccc	gtcaaaaacc	tgaagcgcca	ggatgcccga	1080
atcatcgtgg	gacttttcta	tgagactgaa	gcccggaaa	ttttttgtga	ggtgtacaag	1140
gagcgtctct	ttgggaagaa	gtacgtctgg	ttcctcattg	ggtggtatgc	tgacaattgg	1200
ttcaagatct	acgacccttc	tatcaactgc	acagtggatg	agatgactga	ggcgggtggag	1260
ggccacatca	caactgagat	tgtcatgctg	aatcctgcca	atacccgag	catttccaac	1320
atgacatccc	aggaatttgt	ggagaaacta	accaagcgac	tgaaaagaca	ccctgaggag	1380
acaggaggct	tccaggaggc	accgctggcc	tatgatgcca	tctgggcctt	ggcactggcc	1440
ctgaacaaga	catctggagg	agggcgccgt	tctggtgtgc	gcctggaggga	cttcaactac	1500
aacaaccaga	ccattaccga	ccaaatctac	cgggcaatga	actcttcgtc	ctttgagggg	1560
gtctctggcc	atgtggtgtt	tgatgccagc	ggctctcgga	tggcatggac	gcttatcgag	1620
cagcttcagg	gtggcagcta	caagaagatt	ggctactatg	acagcaccaa	ggatgatctt	1680
tcttggtcca	aaacagataa	atggattgga	gggtccccc	cagctgacca	gacctgggtc	1740
atcaagacat	tccgcttctt	gtcacagaaa	ctctttatct	ccgtctcagt	tctctccagc	1800
ctgggcattg	tctagctgtg	tgtctgtctg	tcctttaaca	tctacaactc	acatgtccgt	1860
tatatccaga	actcacagcc	caacctgaac	aacctgactg	ctgtgggctg	ctcactggct	1920
ttagctgctg	tcttccccct	ggggctcgat	ggttaccaca	ttgggaggaa	ccagtttctt	1980
ttcgtctgcc	aggcccgcct	ctggctcctg	ggcctgggct	ttagtctggg	ctacggttcc	2040

```

atgttcacca agatttggtg ggtccacacg gtcttcacaa agaaggaaga aaagaaggag 2100
tggaaggaaga ctctggaacc ctggaagctg tatgccacag tgggcctgct ggtgggcatg 2160
gatgtcctca ctctcgccat ctggcagatc gtggaccctc tgcaccggac cattgagaca 2220
tttgccaagg aggaacctaa ggaagatatt gacgtctcta ttctgccccca gctggagcat 2280
tgcagctcca ggaagatgaa tacatggctt ggcatcttct atgggttaca ggggctgctg 2340
ctgctgctgg gaatcttcct tgcttatgag accaagagtg tgtccactga gaagatcaat 2400
gatcaccggg ctgtgggcat ggctatctac aatgtggcag tcctgtgcct catcactgct 2460
cctgtcacca tgattctgtc cagccagcag gatgcagcct ttgcctttgc ctctcttgcc 2520
atagttttct cctcctatat cactcttggt gtgctctttg tgcccaagat gcgcaggctg 2580
atcacccgag ggggaatggca gtcggaggcg caggacacca tgaagacagg gtcacgacc 2640
aacaacaacg aggaggagaa gtcccggctg ttggagaagg agaaccgtga actggaaaag 2700
atcattgctg agaaagagga gcgtgtctct gaactgcgcc atcaactcca gtctcggcag 2760
cagctccgct cccggcgcca cccaccgaca cccccagAAC cctctggggg cctgcccagg 2820
ggacccccctg agccccccga ccggcttagc tgtgatggga gtcgagtgcA tttgctttat 2880
aaggcgggcg ccatgactct ggagtccatc atggcgctgct gcctgagcga ggaggccaag 2940
gaagcccggc ggatcaacga cgagatcgag cggcagctcc gcagggacaa gcgggacgcc 3000
cgccgggagc tcaagctgct gctgctcggg acaggagaga gtggcaagag tacgtttatc 3060
aagcagatga gaatcatcca tgggtcagga tactctgatg aagataaaaag gggcttcacc 3120
aagctggtgt atcagaacat cttcacggcc atgcaggcca tgatcagagc catggacaca 3180
ctcaagatcc catacaagta tgagcacaat aaggctcatg cacaattagt tgcagaagtt 3240
gatgtggaga aggtgtctgc ttttgagaat ccatatgtag atgcaataaa gagtttatgg 3300
aatgatcctg gaatccagga atgctatgat agacgacgag aatatcaatt atctgactct 3360
accaaatact atcttaatga cttggaccgc gtactgacc ctgcctacct gcctacgcaa 3420
caagatgtgc ttagagttcg agtccccacc acagggatca tcgaataccc ctttgactta 3480
caaagtgtca ttttcagaat ggtcgatgta gggggccaaa ggtcagagag aagaaaatgg 3540
atacactgct ttgaaaatgt cacctctatc atgtttctag tagcgcttag tgaatatgat 3600
caagttctcg tggagtcaga caatgagaac cgaatggagg aaagcaaggc tctctttaga 3660
acaattatca cataccctg gttccagaac tcctcggtta ttctgttctt aaacaagaaa 3720
gatcttctag aggagaaaat catgtattcc catctagtcg actacttccc agaatatgat 3780
ggaccccaga gagatgcccc ggcagcccga gaattcattc tgaagatggt cgtggacctg 3840
aaccagaca gtgacaaaat tatctactcc cacttcacgt gcgccacaga caccgagaat 3900
atccgctttg tctttgctgc cgtcaaggac accatcctcc agttgaacct gaagggctgc 3960
ggtctgtac

```

<210> 45

<211> 1323

<212> PRT

<213> Artificial Sequence

<220>

<223> Chimeric GABA-BRa1\*Gqo5

<400> 45

```

Met Leu Leu Leu Leu Leu Ala Pro Leu Phe Leu Arg Pro Pro Gly
 1             5             10             15

Ala Gly Gly Ala Gln Thr Pro Asn Ala Thr Ser Glu Gly Cys Gln Ile
 20             25             30

Ile His Pro Pro Trp Glu Gly Gly Ile Arg Tyr Arg Gly Leu Thr Arg
 35             40             45

Asp Gln Val Lys Ala Ile Asn Phe Leu Pro Val Asp Tyr Glu Ile Glu

```

50					55					60					
Tyr	Val	Cys	Arg	Gly	Glu	Arg	Glu	Val	Val	Gly	Pro	Lys	Val	Arg	Lys
65					70					75					80
Cys	Leu	Ala	Asn	Gly	Ser	Trp	Thr	Asp	Met	Asp	Thr	Pro	Ser	Arg	Cys
				85					90					95	
Val	Arg	Ile	Cys	Ser	Lys	Ser	Tyr	Leu	Thr	Leu	Glu	Asn	Gly	Lys	Val
			100					105					110		
Phe	Leu	Thr	Gly	Gly	Asp	Leu	Pro	Ala	Leu	Asp	Gly	Ala	Arg	Val	Asp
			115					120					125		
Phe	Arg	Cys	Asp	Pro	Asp	Phe	His	Leu	Val	Gly	Ser	Ser	Arg	Ser	Ile
			130					135					140		
Cys	Ser	Gln	Gly	Gln	Trp	Ser	Thr	Pro	Lys	Pro	His	Cys	Gln	Val	Asn
145															160
Arg	Thr	Pro	His	Ser	Glu	Arg	Arg	Ala	Val	Tyr	Ile	Gly	Ala	Leu	Phe
				165					170						175
Pro	Met	Ser	Gly	Gly	Trp	Pro	Gly	Gly	Gln	Ala	Cys	Gln	Pro	Ala	Val
			180					185						190	
Glu	Met	Ala	Leu	Glu	Asp	Val	Asn	Ser	Arg	Arg	Asp	Ile	Leu	Pro	Asp
			195				200					205			
Tyr	Glu	Leu	Lys	Leu	Ile	His	His	Asp	Ser	Lys	Cys	Asp	Pro	Gly	Gln
			210				215					220			
Ala	Thr	Lys	Tyr	Leu	Tyr	Glu	Leu	Leu	Tyr	Asn	Asp	Pro	Ile	Lys	Ile
225							230					235			240
Ile	Leu	Met	Pro	Gly	Cys	Ser	Ser	Val	Ser	Thr	Leu	Val	Ala	Glu	Ala
				245					250					255	
Ala	Arg	Met	Trp	Asn	Leu	Ile	Val	Leu	Ser	Tyr	Gly	Ser	Ser	Ser	Pro
			260					265					270		
Ala	Leu	Ser	Asn	Arg	Gln	Arg	Phe	Pro	Thr	Phe	Phe	Arg	Thr	His	Pro
			275				280					285			
Ser	Ala	Thr	Leu	His	Asn	Pro	Thr	Arg	Val	Lys	Leu	Phe	Glu	Lys	Trp
			290				295					300			
Gly	Trp	Lys	Lys	Ile	Ala	Thr	Ile	Gln	Gln	Thr	Thr	Glu	Val	Phe	Thr
305							310					315			320
Ser	Thr	Leu	Asp	Asp	Leu	Glu	Glu	Arg	Val	Lys	Glu	Ala	Gly	Ile	Glu
				325					330					335	
Ile	Thr	Phe	Arg	Gln	Ser	Phe	Phe	Ser	Asp	Pro	Ala	Val	Pro	Val	Lys
				340				345					350		



Asn Leu Lys Arg Gln Asp Ala Arg Ile Ile Val Gly Leu Phe Tyr Glu  
 355 360 365  
 Thr Glu Ala Arg Lys Val Phe Cys Glu Val Tyr Lys Glu Arg Leu Phe  
 370 375 380  
 Gly Lys Lys Tyr Val Trp Phe Leu Ile Gly Trp Tyr Ala Asp Asn Trp  
 385 390 395 400  
 Phe Lys Ile Tyr Asp Pro Ser Ile Asn Cys Thr Val Asp Glu Met Thr  
 405 410 415  
 Glu Ala Val Glu Gly His Ile Thr Thr Glu Ile Val Met Leu Asn Pro  
 420 425 430  
 Ala Asn Thr Arg Ser Ile Ser Asn Met Thr Ser Gln Glu Phe Val Glu  
 435 440 445  
 Lys Leu Thr Lys Arg Leu Lys Arg His Pro Glu Glu Thr Gly Gly Phe  
 450 455 460  
 Gln Glu Ala Pro Leu Ala Tyr Asp Ala Ile Trp Ala Leu Ala Leu Ala  
 465 470 475 480  
 Leu Asn Lys Thr Ser Gly Gly Gly Gly Arg Ser Gly Val Arg Leu Glu  
 485 490 495  
 Asp Phe Asn Tyr Asn Asn Gln Thr Ile Thr Asp Gln Ile Tyr Arg Ala  
 500 505 510  
 Met Asn Ser Ser Ser Phe Glu Gly Val Ser Gly His Val Val Phe Asp  
 515 520 525  
 Ala Ser Gly Ser Arg Met Ala Trp Thr Leu Ile Glu Gln Leu Gln Gly  
 530 535 540  
 Gly Ser Tyr Lys Lys Ile Gly Tyr Tyr Asp Ser Thr Lys Asp Asp Leu  
 545 550 555 560  
 Ser Trp Ser Lys Thr Asp Lys Trp Ile Gly Gly Ser Pro Pro Ala Asp  
 565 570 575  
 Gln Thr Leu Val Ile Lys Thr Phe Arg Phe Leu Ser Gln Lys Leu Phe  
 580 585 590  
 Ile Ser Val Ser Val Leu Ser Ser Leu Gly Ile Val Leu Ala Val Val  
 595 600 605  
 Cys Leu Ser Phe Asn Ile Tyr Asn Ser His Val Arg Tyr Ile Gln Asn  
 610 615 620  
 Ser Gln Pro Asn Leu Asn Asn Leu Thr Ala Val Gly Cys Ser Leu Ala  
 625 630 635 640  
 Leu Ala Ala Val Phe Pro Leu Gly Leu Asp Gly Tyr His Ile Gly Arg  
 645 650 655

Asn Gln Phe Pro Phe Val Cys Gln Ala Arg Leu Trp Leu Leu Gly Leu  
 660 665 670  
 Gly Phe Ser Leu Gly Tyr Gly Ser Met Phe Thr Lys Ile Trp Trp Val  
 675 680 685  
 His Thr Val Phe Thr Lys Lys Glu Glu Lys Lys Glu Trp Arg Lys Thr  
 690 695 700  
 Leu Glu Pro Trp Lys Leu Tyr Ala Thr Val Gly Leu Leu Val Gly Met  
 705 710 715 720  
 Asp Val Leu Thr Leu Ala Ile Trp Gln Ile Val Asp Pro Leu His Arg  
 725 730 735  
 Thr Ile Glu Thr Phe Ala Lys Glu Glu Pro Lys Glu Asp Ile Asp Val  
 740 745 750  
 Ser Ile Leu Pro Gln Leu Glu His Cys Ser Ser Arg Lys Met Asn Thr  
 755 760 765  
 Trp Leu Gly Ile Phe Tyr Gly Tyr Lys Gly Leu Leu Leu Leu Leu Gly  
 770 775 780  
 Ile Phe Leu Ala Tyr Glu Thr Lys Ser Val Ser Thr Glu Lys Ile Asn  
 785 790 795 800  
 Asp His Arg Ala Val Gly Met Ala Ile Tyr Asn Val Ala Val Leu Cys  
 805 810 815  
 Leu Ile Thr Ala Pro Val Thr Met Ile Leu Ser Ser Gln Gln Asp Ala  
 820 825 830  
 Ala Phe Ala Phe Ala Ser Leu Ala Ile Val Phe Ser Ser Tyr Ile Thr  
 835 840 845  
 Leu Val Val Leu Phe Val Pro Lys Met Arg Arg Leu Ile Thr Arg Gly  
 850 855 860  
 Glu Trp Gln Ser Glu Ala Gln Asp Thr Met Lys Thr Gly Ser Ser Thr  
 865 870 875 880  
 Asn Asn Asn Glu Glu Glu Lys Ser Arg Leu Leu Glu Lys Glu Asn Arg  
 885 890 895  
 Glu Leu Glu Lys Ile Ile Ala Glu Lys Glu Glu Arg Val Ser Glu Leu  
 900 905 910  
 Arg His Gln Leu Gln Ser Arg Gln Gln Leu Arg Ser Arg Arg His Pro  
 915 920 925  
 Pro Thr Pro Pro Glu Pro Ser Gly Gly Leu Pro Arg Gly Pro Pro Glu  
 930 935 940  
 Pro Pro Asp Arg Leu Ser Cys Asp Gly Ser Arg Val His Leu Leu Tyr

945	950	955	960
Lys Ala Ala Ala Met Thr Leu Glu Ser Ile Met Ala Cys Cys Leu Ser			
965		970	975
Glu Glu Ala Lys Glu Ala Arg Arg Ile Asn Asp Glu Ile Glu Arg Gln			
980		985	990
Leu Arg Arg Asp Lys Arg Asp Ala Arg Arg Glu Leu Lys Leu Leu Leu			
995		1000	1005
Leu Gly Thr Gly Glu Ser Gly Lys Ser Thr Phe Ile Lys Gln Met Arg			
1010		1015	1020
Ile Ile His Gly Ser Gly Tyr Ser Asp Glu Asp Lys Arg Gly Phe Thr			
1025		1030	1035
Lys Leu Val Tyr Gln Asn Ile Phe Thr Ala Met Gln Ala Met Ile Arg			
		1045	1050
Ala Met Asp Thr Leu Lys Ile Pro Tyr Lys Tyr Glu His Asn Lys Ala			
		1060	1065
His Ala Gln Leu Val Arg Glu Val Asp Val Glu Lys Val Ser Ala Phe			
		1075	1080
Glu Asn Pro Tyr Val Asp Ala Ile Lys Ser Leu Trp Asn Asp Pro Gly			
		1090	1095
Ile Gln Glu Cys Tyr Asp Arg Arg Arg Glu Tyr Gln Leu Ser Asp Ser			
1105		1110	1115
Thr Lys Tyr Tyr Leu Asn Asp Leu Asp Arg Val Ala Asp Pro Ala Tyr			
		1125	1130
Leu Pro Thr Gln Gln Asp Val Leu Arg Val Arg Val Pro Thr Thr Gly			
		1140	1145
Ile Ile Glu Tyr Pro Phe Asp Leu Gln Ser Val Ile Phe Arg Met Val			
		1155	1160
Asp Val Gly Gly Gln Arg Ser Glu Arg Arg Lys Trp Ile His Cys Phe			
		1170	1175
Glu Asn Val Thr Ser Ile Met Phe Leu Val Ala Leu Ser Glu Tyr Asp			
1185		1190	1195
Gln Val Leu Val Glu Ser Asp Asn Glu Asn Arg Met Glu Glu Ser Lys			
		1205	1210
Ala Leu Phe Arg Thr Ile Ile Thr Tyr Pro Trp Phe Gln Asn Ser Ser			
		1220	1225
Val Ile Leu Phe Leu Asn Lys Lys Asp Leu Leu Glu Glu Lys Ile Met			
		1235	1240
			1245

Tyr Ser His Leu Val Asp Tyr Phe Pro Glu Tyr Asp Gly Pro Gln Arg  
 1250 1255 1260  
 Asp Ala Gln Ala Ala Arg Glu Phe Ile Leu Lys Met Phe Val Asp Leu  
 1265 1270 1275 1280  
 Asn Pro Asp Ser Asp Lys Ile Ile Tyr Ser His Phe Thr Cys Ala Thr  
 1285 1290 1295  
 Asp Thr Glu Asn Ile Arg Phe Val Phe Ala Ala Val Lys Asp Thr Ile  
 1300 1305 1310  
 Leu Gln Leu Asn Leu Lys Gly Cys Gly Leu Tyr  
 1315 1320

<210> 46  
 <211> 4231  
 <212> DNA  
 <213> Artificial Sequence  
 <220>  
 <223> Chimeric pmGluR2//CaR\*G(qi5+3Ala  
 <400> 46

atgggatcgc tgccttgcgct cccggcactg ctgctgctgt ggggtgctgt ggctgagggc 60  
 ccagccaaga aggtgctgac cctggagggg gacttggtgc tgggtgggct gttcccagtg 120  
 caccagaagg gcggcccagc agaggactgt ggtcctgtca atgagcaccg tggcatccag 180  
 cgcctggagg ccatgctttt tgcactggac cgcatacaacc gtgaccgca cctgctgcct 240  
 ggcgtgcgcc tgggtgcaca catcctcgac agttgctcca aggacacaca tgcgtggag 300  
 caggcactgg actttgtgcg tgcctcactc agcctgggtg ctgatggctc acgccacatc 360  
 tgccccgacg gctcttatgc gacccatggt gatgctccca ctgccatcac tgggtgttatt 420  
 ggcggttcct acagtgatgt ctccatccag gtggccaacc tcttgaggct atttcagatc 480  
 ccacagatta gctacgcctc taccagtgcc aagctgagtg acaagtcccg ctatgactac 540  
 tttgcccga cagtgcctcc tgacttcttc caagccaagg ccatggctga gattctccgc 600  
 ttcttcaact ggacctatgt gtccactgtg gcgtctgagg gcgactatgg cgagacaggc 660  
 attgaggcct ttgagctaga ggctcgtgcc cgcaacatct gtgtggccac ctcgagaaa 720  
 gtgggcccgt ccatgagccg cgcggccttt gaggggtgtg tgcgagccct gctgcagaag 780  
 cccagtgcct gcgtggtgtg cctgttcacc cgttctgagg atgcccggga gctgcttgtc 840  
 gccagccagc gcctcaatgc cagcttcacc tgggtggcca gtgatggtt gggggccctg 900  
 gagagtgtgg tggcaggcag tgagggggct gctgaggggt ctatcaccat cgagctggcc 960  
 tcctacccca tcagtgaact tgctcctac ttccagagcc tggacccttg gaacaacagc 1020  
 cggaaccctt ggttcogtga attctgggag cagaggttcc gctgcagctt ccggcagcga 1080  
 gactgcgcag cccactctct ccgggctgtg ccttttgagc aggagtccaa gatcatgttt 1140  
 gtggtcaatg cagtgtacgc catggcccat gcgctccaca acatgcaccg tgccctctgc 1200  
 cccaacacca cccggctctg tgacgcgatg cggccagtta acgggcgccc cctctacaag 1260  
 gactttgtgc tcaacgtcaa gtttgatgcc ccttttcgcc cagctgacac ccacaatgag 1320  
 gtccgctttg accgcttttg tgatggtatt ggccgctaca acatcttcac ctatctgcgt 1380  
 gcaggcagtg ggcgctatcg ctaccagaag gtgggctact gggcagaagg cttgactctg 1440  
 gacaccagcc tcatcccatg ggctcacc tcagccggcc cctgcccgc ctctcgctgc 1500  
 agtgagccct gcctccagaa tgaggtgaag agtgtgcagc cgggcgaagt ctgctgctgg 1560  
 ctctgcattc cgtgccagcc ctatgagtag cgattggacg aattcacttg cgctgattgt 1620  
 ggcttgggct actggcccaa tgccagcctg actggctgct tcgaactgcc ccaggagtac 1680  
 atccgctggg gcgatgcctg ggctgtggga cctgtcacca tcgcctgcct cggtgccctg 1740  
 gccaccctct ttgtgctggg tgtctttgtg cggcacaatg ccacaccagt ggtcaaggcc 1800

```

tcaggctcggg agctctgcta catcctgctg ggtggtgtct tcctctgcta ctgcatgacc 1860
ttcatcttca ttgccaagcc atccacggca gtgtgtacct tacggcgtct tggtttgggc 1920
actgccttct ctgtctgcta ctcagccctg ctcaccaaga ccaaccgcat tgcacgcatc 1980
ttcggtgggg cccgggaggg tgcccagcgg ccacgcttca tcagtcctgc ctcacagggtg 2040
gccatctgcc tggcacttat ctggggccag ctgctcatcg tggtcgcctg gctggtggtg 2100
gaggcaccgg gcacaggcaa ggagacagcc cccgaacggc gggagggtgg gacactgcgc 2160
tgcaaccacc gcgatgcaag tatgttgggc tcgctggcct acaatgtgct cctcatcgcg 2220
ctctgcacgc tttatgcctt caagactcgc aagtgcctcg aaaacttcaa cgaggccaag 2280
ttcattggct tcacatgta caccactgc atcatctggc tggcattcct gccatcttc 2340
tatgtcacct ccagtgaacta ccgggtacag accaccacca tgtgcgtgtc agtcagcctc 2400
agcggctccg tgggtgcttg ctgcctcttt gcgccaagc tgcacatcat cctcttccag 2460
ccgcagaaga acaccatcga ggagggtgct tgcagcaccg cagctcacgc tttcaagggtg 2520
gctgcccggg ccacgctgcg ccgcagcaac gtctcccgca agcgggtccag cagccttgga 2580
ggctccacgg gatccacccc ctctctctcc atcagcagca agagcaacag cgaagacca 2640
ttcccacagc ccgagaggca gaagcagcag cagccgctgg ccctaacca gcaagagcag 2700
cagcagcagc cctgaccct cccacagcag caacgatctc agcagcagcc cagatgcaag 2760
cagaaggta tctttggcag cggcacggtc accttctcac tgagctttga tgagcctcag 2820
aagaacgcca tggcccacgg gaattctacg caccagaact ccctggaggc ccagaaaagc 2880
agcgatcgc tgaccgcaca ccagccatta ctcccgtgc agtgcgggga aacggactta 2940
gatctgaccg tccaggaaac aggtctgcaa ggacctgtgg gtggagacca gcggccagag 3000
gtggaggacc ctgaagagtt gtcccagca cttgtagtgt ccagttcaca gagctttgtc 3060
atcagtgggtg gaggcagcac tgttacagaa aacgtagtga attcagcggc cgccatgact 3120
ctggagtcca tcatggcgtg ctgcctgagc gaggaggcca aggaagcccg gcggatcaac 3180
gacgagatcg agcggcagct ccgcagggac aagcgggacg cccgccggga gctcaagctg 3240
ctgctgctcg ggacaggaga gagtggcaag agtacgttta tcaagcagat gagaatcatc 3300
catgggtcag gatactctga tgaagataaa aggggcttca ccaagctggt gtatcagaac 3360
atcttcacgg ccatgcaggc catgatcaga gccatggaca cactcaagat cccatacaag 3420
tatgagcaca ataaggctca tgcacaatta gttcgagaag ttgatgtgga gaaggtgtct 3480
gcttttgaga atccatatgt agatgcaata aagagtttat ggaatgatcc tggaatccag 3540
gaatgctatg atagcagcg agaatatcaa ttatctgact ctaccaaata ctatcttaat 3600
gacttggacc gcgtagctga ccctgcctac ctgcctacgc aacaagatgt gcttagagtt 3660
cgagtcccca ccacagggat catcgaatac ccctttgact taaaaagtgt cattttcaga 3720
atgggtcgatg tagggggcca aaggctcagag agaagaaaat ggatacactg ctttgaaaat 3780
gtcacctcta tcatgtttct agtagcgctt agtgaatatg atcaagttct cgtggagtca 3840
gacaatgaga accgaatgga ggaaagcaag gctctcttta gaacaattat cacatacccc 3900
tggttccaga actcctcggg tattctgttc ttaaacaaga aagatcttct agaggagaaa 3960
atcatgtatt cccatctagt cgactacttc ccagaatatg atggacccca gagagatgcc 4020
caggcagccc gagaattcat tctgaagatg ttcgtggacc tgaaccaga cagtgacaaa 4080
attatctact cccacttcac gtgcgccaca gacaccgaga atatccgctt tgtctttgct 4140
gccgtcaagg acaccatcct ccagttgaac ctgaaggact gcggtctggt ctaattgtgc 4200
ctcctagaca cccgccctgc ccttccctgg t 4231

```

<210> 47

<211> 1397

<212> PRT

<213> Artificial Sequence

<220>

<223> Chimeric pmGluR2//CaR\*G(qi5+3Ala

<400> 47

Met Gly Ser Leu Leu Ala Leu Pro Ala Leu Leu Leu Leu Trp Gly Ala  
1 5 10 15

Val Ala Glu Gly Pro Ala Lys Lys Val Leu Thr Leu Glu Gly Asp Leu  
 20 25 30  
 Val Leu Gly Gly Leu Phe Pro Val His Gln Lys Gly Gly Pro Ala Glu  
 35 40 45  
 Asp Cys Gly Pro Val Asn Glu His Arg Gly Ile Gln Arg Leu Glu Ala  
 50 55 60  
 Met Leu Phe Ala Leu Asp Arg Ile Asn Arg Asp Pro His Leu Leu Pro  
 65 70 75 80  
 Gly Val Arg Leu Gly Ala His Ile Leu Asp Ser Cys Ser Lys Asp Thr  
 85 90 95  
 His Ala Leu Glu Gln Ala Leu Asp Phe Val Arg Ala Ser Leu Ser Arg  
 100 105 110  
 Gly Ala Asp Gly Ser Arg His Ile Cys Pro Asp Gly Ser Tyr Ala Thr  
 115 120 125  
 His Gly Asp Ala Pro Thr Ala Ile Thr Gly Val Ile Gly Gly Ser Tyr  
 130 135 140  
 Ser Asp Val Ser Ile Gln Val Ala Asn Leu Leu Arg Leu Phe Gln Ile  
 145 150 155 160  
 Pro Gln Ile Ser Tyr Ala Ser Thr Ser Ala Lys Leu Ser Asp Lys Ser  
 165 170 175  
 Arg Tyr Asp Tyr Phe Ala Arg Thr Val Pro Pro Asp Phe Phe Gln Ala  
 180 185 190  
 Lys Ala Met Ala Glu Ile Leu Arg Phe Phe Asn Trp Thr Tyr Val Ser  
 195 200 205  
 Thr Val Ala Ser Glu Gly Asp Tyr Gly Glu Thr Gly Ile Glu Ala Phe  
 210 215 220  
 Glu Leu Glu Ala Arg Ala Arg Asn Ile Cys Val Ala Thr Ser Glu Lys  
 225 230 235 240  
 Val Gly Arg Ala Met Ser Arg Ala Ala Phe Glu Gly Val Val Arg Ala  
 245 250 255  
 Leu Leu Gln Lys Pro Ser Ala Arg Val Ala Val Leu Phe Thr Arg Ser  
 260 265 270  
 Glu Asp Ala Arg Glu Leu Leu Ala Ala Ser Gln Arg Leu Asn Ala Ser  
 275 280 285  
 Phe Thr Trp Val Ala Ser Asp Gly Trp Gly Ala Leu Glu Ser Val Val  
 290 295 300  
 Ala Gly Ser Glu Gly Ala Ala Glu Gly Ala Ile Thr Ile Glu Leu Ala

305		310		315		320
Ser Tyr Pro Ile	Ser Asp Phe Ala Ser Tyr Phe Gln Ser Leu Asp Pro					
	325		330		335	
Trp Asn Asn Ser Arg Asn Pro Trp Phe Arg Glu Phe Trp Glu Gln Arg						
	340		345		350	
Phe Arg Cys Ser Phe Arg Gln Arg Asp Cys Ala Ala His Ser Leu Arg						
	355		360		365	
Ala Val Pro Phe Glu Gln Glu Ser Lys Ile Met Phe Val Val Asn Ala						
	370		375		380	
Val Tyr Ala Met Ala His Ala Leu His Asn Met His Arg Ala Leu Cys						
	385		390		395	400
Pro Asn Thr Thr Arg Leu Cys Asp Ala Met Arg Pro Val Asn Gly Arg						
		405		410		415
Arg Leu Tyr Lys Asp Phe Val Leu Asn Val Lys Phe Asp Ala Pro Phe						
		420		425		430
Arg Pro Ala Asp Thr His Asn Glu Val Arg Phe Asp Arg Phe Gly Asp						
		435		440		445
Gly Ile Gly Arg Tyr Asn Ile Phe Thr Tyr Leu Arg Ala Gly Ser Gly						
		450		455		460
Arg Tyr Arg Tyr Gln Lys Val Gly Tyr Trp Ala Glu Gly Leu Thr Leu						
		465		470		475
Asp Thr Ser Leu Ile Pro Trp Ala Ser Pro Ser Ala Gly Pro Leu Pro						
		485		490		495
Ala Ser Arg Cys Ser Glu Pro Cys Leu Gln Asn Glu Val Lys Ser Val						
		500		505		510
Gln Pro Gly Glu Val Cys Cys Trp Leu Cys Ile Pro Cys Gln Pro Tyr						
		515		520		525
Glu Tyr Arg Leu Asp Glu Phe Thr Cys Ala Asp Cys Gly Leu Gly Tyr						
		530		535		540
Trp Pro Asn Ala Ser Leu Thr Gly Cys Phe Glu Leu Pro Gln Glu Tyr						
		545		550		555
Ile Arg Trp Gly Asp Ala Trp Ala Val Gly Pro Val Thr Ile Ala Cys						
		565		570		575
Leu Gly Ala Leu Ala Thr Leu Phe Val Leu Gly Val Phe Val Arg His						
		580		585		590
Asn Ala Thr Pro Val Val Lys Ala Ser Gly Arg Glu Leu Cys Tyr Ile						
		595		600		605

Leu Leu Gly Gly Val Phe Leu Cys Tyr Cys Met Thr Phe Ile Phe Ile  
 610 615 620  
 Ala Lys Pro Ser Thr Ala Val Cys Thr Leu Arg Arg Leu Gly Leu Gly  
 625 630 635 640  
 Thr Ala Phe Ser Val Cys Tyr Ser Ala Leu Leu Thr Lys Thr Asn Arg  
 645 650 655  
 Ile Ala Arg Ile Phe Gly Gly Ala Arg Glu Gly Ala Gln Arg Pro Arg  
 660 665 670  
 Phe Ile Ser Pro Ala Ser Gln Val Ala Ile Cys Leu Ala Leu Ile Ser  
 675 680 685  
 Gly Gln Leu Leu Ile Val Val Ala Trp Leu Val Val Glu Ala Pro Gly  
 690 695 700  
 Thr Gly Lys Glu Thr Ala Pro Glu Arg Arg Glu Val Val Thr Leu Arg  
 705 710 715 720  
 Cys Asn His Arg Asp Ala Ser Met Leu Gly Ser Leu Ala Tyr Asn Val  
 725 730 735  
 Leu Leu Ile Ala Leu Cys Thr Leu Tyr Ala Phe Lys Thr Arg Lys Cys  
 740 745 750  
 Pro Glu Asn Phe Asn Glu Ala Lys Phe Ile Gly Phe Thr Met Tyr Thr  
 755 760 765  
 Thr Cys Ile Ile Trp Leu Ala Phe Leu Pro Ile Phe Tyr Val Thr Ser  
 770 775 780  
 Ser Asp Tyr Arg Val Gln Thr Thr Thr Met Cys Val Ser Val Ser Leu  
 785 790 795 800  
 Ser Gly Ser Val Val Leu Gly Cys Leu Phe Ala Pro Lys Leu His Ile  
 805 810 815  
 Ile Leu Phe Gln Pro Gln Lys Asn Thr Ile Glu Glu Val Arg Cys Ser  
 820 825 830  
 Thr Ala Ala His Ala Phe Lys Val Ala Ala Arg Ala Thr Leu Arg Arg  
 835 840 845  
 Ser Asn Val Ser Arg Lys Arg Ser Ser Ser Leu Gly Gly Ser Thr Gly  
 850 855 860  
 Ser Thr Pro Ser Ser Ser Ile Ser Ser Lys Ser Asn Ser Glu Asp Pro  
 865 870 875 880  
 Phe Pro Gln Pro Glu Arg Gln Lys Gln Gln Gln Pro Leu Ala Leu Thr  
 885 890 895  
 Gln Gln Glu Gln Gln Gln Gln Pro Leu Thr Leu Pro Gln Gln Gln Arg  
 900 905 910



Asp Leu Asp Arg Val Ala Asp Pro Ala Tyr Leu Pro Thr Gln Gln Asp

1205										1210										1215																																			
Val	Leu	Arg	Val	Arg	Val	Pro	Thr	Thr	Gly	Ile	Ile	Glu	Tyr	Pro	Phe					Val	Leu	Arg	Val	Arg	Val	Pro	Thr	Thr	Gly	Ile	Ile	Glu	Tyr	Pro	Phe			Val	Leu	Arg	Val	Arg	Val	Pro	Thr	Thr	Gly	Ile	Ile	Glu	Tyr	Pro	Phe		
Asp	Leu	Gln	Ser	Val	Ile	Phe	Arg	Met	Val	Asp	Val	Gly	Gly	Gln	Arg					Asp	Leu	Gln	Ser	Val	Ile	Phe	Arg	Met	Val	Asp	Val	Gly	Gly	Gln	Arg			Asp	Leu	Gln	Ser	Val	Ile	Phe	Arg	Met	Val	Asp	Val	Gly	Gly	Gln	Arg		
Ser	Glu	Arg	Arg	Lys	Trp	Ile	His	Cys	Phe	Glu	Asn	Val	Thr	Ser	Ile					Ser	Glu	Arg	Arg	Lys	Trp	Ile	His	Cys	Phe	Glu	Asn	Val	Thr	Ser	Ile			Ser	Glu	Arg	Arg	Lys	Trp	Ile	His	Cys	Phe	Glu	Asn	Val	Thr	Ser	Ile		
Met	Phe	Leu	Val	Ala	Leu	Ser	Glu	Tyr	Asp	Gln	Val	Leu	Val	Glu	Ser					Met	Phe	Leu	Val	Ala	Leu	Ser	Glu	Tyr	Asp	Gln	Val	Leu	Val	Glu	Ser			Met	Phe	Leu	Val	Ala	Leu	Ser	Glu	Tyr	Asp	Gln	Val	Leu	Val	Glu	Ser		
Asp	Asn	Glu	Asn	Arg	Met	Glu	Glu	Ser	Lys	Ala	Leu	Phe	Arg	Thr	Ile					Asp	Asn	Glu	Asn	Arg	Met	Glu	Glu	Ser	Lys	Ala	Leu	Phe	Arg	Thr	Ile			Asp	Asn	Glu	Asn	Arg	Met	Glu	Glu	Ser	Lys	Ala	Leu	Phe	Arg	Thr	Ile		
Ile	Thr	Tyr	Pro	Trp	Phe	Gln	Asn	Ser	Ser	Val	Ile	Leu	Phe	Leu	Asn					Ile	Thr	Tyr	Pro	Trp	Phe	Gln	Asn	Ser	Ser	Val	Ile	Leu	Phe	Leu	Asn			Ile	Thr	Tyr	Pro	Trp	Phe	Gln	Asn	Ser	Ser	Val	Ile	Leu	Phe	Leu	Asn		
Lys	Lys	Asp	Leu	Leu	Glu	Glu	Lys	Ile	Met	Tyr	Ser	His	Leu	Val	Asp					Lys	Lys	Asp	Leu	Leu	Glu	Glu	Lys	Ile	Met	Tyr	Ser	His	Leu	Val	Asp			Lys	Lys	Asp	Leu	Leu	Glu	Glu	Lys	Ile	Met	Tyr	Ser	His	Leu	Val	Asp		
Tyr	Phe	Pro	Glu	Tyr	Asp	Gly	Pro	Gln	Arg	Asp	Ala	Gln	Ala	Ala	Arg					Tyr	Phe	Pro	Glu	Tyr	Asp	Gly	Pro	Gln	Arg	Asp	Ala	Gln	Ala	Ala	Arg			Tyr	Phe	Pro	Glu	Tyr	Asp	Gly	Pro	Gln	Arg	Asp	Ala	Gln	Ala	Ala	Arg		
Glu	Phe	Ile	Leu	Lys	Met	Phe	Val	Asp	Leu	Asn	Pro	Asp	Ser	Asp	Lys					Glu	Phe	Ile	Leu	Lys	Met	Phe	Val	Asp	Leu	Asn	Pro	Asp	Ser	Asp	Lys			Glu	Phe	Ile	Leu	Lys	Met	Phe	Val	Asp	Leu	Asn	Pro	Asp	Ser	Asp	Lys		
Ile	Ile	Tyr	Ser	His	Phe	Thr	Cys	Ala	Thr	Asp	Thr	Glu	Asn	Ile	Arg					Ile	Ile	Tyr	Ser	His	Phe	Thr	Cys	Ala	Thr	Asp	Thr	Glu	Asn	Ile	Arg			Ile	Ile	Tyr	Ser	His	Phe	Thr	Cys	Ala	Thr	Asp	Thr	Glu	Asn	Ile	Arg		
Phe	Val	Phe	Ala	Ala	Val	Lys	Asp	Thr	Ile	Leu	Gln	Leu	Asn	Leu	Lys					Phe	Val	Phe	Ala	Ala	Val	Lys	Asp	Thr	Ile	Leu	Gln	Leu	Asn	Leu	Lys			Phe	Val	Phe	Ala	Ala	Val	Lys	Asp	Thr	Ile	Leu	Gln	Leu	Asn	Leu	Lys		
Asp	Cys	Gly	Leu	Phe																Asp	Cys	Gly	Leu	Phe														Asp	Cys	Gly	Leu	Phe													

&lt;210&gt; 48

&lt;211&gt; 905

&lt;212&gt; PRT

&lt;213&gt; Chimeric construct ph8SPmGluR4

&lt;400&gt; 48

Met	Val	Cys	Glu	Gly	Lys	Arg	Ser	Ala	Ser	Cys	Pro	Cys	Phe	Phe	Leu					Met	Val	Cys	Glu	Gly	Lys	Arg	Ser	Ala	Ser	Cys	Pro	Cys	Phe	Phe	Leu			Met	Val	Cys	Glu	Gly	Lys	Arg	Ser	Ala	Ser	Cys	Pro	Cys	Phe	Phe	Leu		
1					5				10					15					1					5				10			15					1					5				10				15						
Leu	Thr	Ala	Lys	Phe	Tyr	Trp	Ile	Leu	Thr	Met	Met	Gln	Arg	Thr	His				Leu	Thr	Ala	Lys	Phe	Tyr	Trp	Ile	Leu	Thr	Met	Met	Gln	Arg	Thr	His			Leu	Thr	Ala	Lys	Phe	Tyr	Trp	Ile	Leu	Thr	Met	Met	Gln	Arg	Thr	His			
			20				25					30										20				25				30									20				25					30							
Ser	Gln	Glu	Tyr	Ala	His	Ser	Ile	Arg	Ile	Asp	Gly	Asp	Ile	Thr	Leu				Ser	Gln	Glu	Tyr	Ala	His	Ser	Ile	Arg	Ile	Asp	Gly	Asp	Ile	Thr	Leu			Ser	Gln	Glu	Tyr	Ala	His	Ser	Ile	Arg	Ile	Asp	Gly	Asp	Ile	Thr	Leu			
			35			40				45												35			40				45										35			40					45								
Gly	Gly	Leu	Phe	Pro	Val	His	Gly	Arg	Gly	Ser	Glu	Gly	Lys	Pro	Cys				Gly	Gly	Leu	Phe	Pro	Val	His	Gly	Arg	Gly	Ser	Glu	Gly	Lys	Pro	Cys			Gly	Gly	Leu	Phe	Pro	Val	His	Gly	Arg	Gly	Ser	Glu	Gly	Lys	Pro	Cys			
			50			55				60												50			55				60										50			55					60								
Gly	Glu	Leu	Lys	Lys	Glu	Lys	Gly	Ile	His	Arg	Leu	Glu	Ala	Met	Leu				Gly	Glu	Leu	Lys	Lys	Glu	Lys	Gly	Ile	His	Arg	Leu	Glu	Ala	Met	Leu			Gly	Glu	Leu	Lys	Lys	Glu	Lys	Gly	Ile	His	Arg	Leu	Glu	Ala	Met	Leu			
65					70				75					80					65					70				75				80				65					70				75				80						

Phe Ala Leu Asp Arg Ile Asn Asn Asp Pro Asp Leu Leu Pro Asn Ile  
 85 90 95  
 Thr Leu Gly Ala Arg Ile Leu Asp Thr Cys Ser Arg Asp Thr His Ala  
 100 105 110  
 Leu Glu Gln Ser Leu Thr Phe Val Gln Ala Leu Ile Glu Lys Asp Gly  
 115 120 125  
 Thr Glu Val Arg Cys Gly Ser Gly Gly Pro Pro Ile Ile Thr Lys Pro  
 130 135 140  
 Glu Arg Val Val Gly Val Ile Gly Ala Ser Gly Ser Ser Val Ser Ile  
 145 150 155 160  
 Met Val Ala Asn Ile Leu Arg Leu Phe Lys Ile Pro Gln Ile Ser Tyr  
 165 170 175  
 Ala Ser Thr Ala Pro Asp Leu Ser Asp Asn Ser Arg Tyr Asp Phe Phe  
 180 185 190  
 Ser Arg Val Val Pro Ser Asp Thr Tyr Gln Ala Gln Ala Met Val Asp  
 195 200 205  
 Ile Val Arg Ala Leu Lys Trp Asn Tyr Val Ser Thr Val Ala Ser Glu  
 210 215 220  
 Gly Ser Tyr Gly Glu Ser Gly Val Glu Ala Phe Ile Gln Lys Ser Arg  
 225 230 235 240  
 Glu Asp Gly Gly Val Cys Ile Ala Gln Ser Val Lys Ile Pro Arg Glu  
 245 250 255  
 Pro Lys Ala Gly Glu Phe Asp Lys Ile Ile Arg Arg Leu Leu Glu Thr  
 260 265 270  
 Ser Asn Ala Arg Ala Val Ile Ile Phe Ala Asn Glu Asp Asp Ile Arg  
 275 280 285  
 Arg Val Leu Glu Ala Ala Arg Arg Ala Asn Gln Thr Gly His Phe Phe  
 290 295 300  
 Trp Met Gly Ser Asp Ser Trp Gly Ser Lys Ile Ala Pro Val Leu His  
 305 310 315 320  
 Leu Glu Glu Val Ala Glu Gly Ala Val Thr Ile Leu Pro Lys Arg Met  
 325 330 335  
 Ser Val Arg Gly Phe Asp Arg Tyr Phe Ser Ser Arg Thr Leu Asp Asn  
 340 345 350  
 Asn Arg Arg Asn Ile Trp Phe Ala Glu Phe Trp Glu Asp Asn Phe His  
 355 360 365  
 Cys Lys Leu Ser Arg His Ala Leu Lys Lys Gly Ser His Val Lys Lys  
 370 375 380  
 Cys Thr Asn Arg Glu Arg Ile Gly Gln Asp Ser Ala Tyr Glu Gln Glu  
 385 390 395 400  
 Gly Lys Val Gln Phe Val Ile Asp Ala Val Tyr Ala Met Gly His Ala  
 405 410 415  
 Leu His Ala Met His Arg Asp Leu Cys Pro Gly Arg Val Gly Leu Cys  
 420 425 430  
 Pro Arg Met Asp Pro Val Asp Gly Thr Gln Leu Leu Lys Tyr Ile Arg  
 435 440 445  
 Asn Val Asn Phe Ser Gly Ile Ala Gly Asn Pro Val Thr Phe Asn Glu  
 450 455 460  
 Asn Gly Asp Ala Pro Gly Arg Tyr Asp Ile Tyr Gln Tyr Gln Leu Arg  
 465 470 475 480  
 Asn Asp Ser Ala Glu Tyr Lys Val Ile Gly Ser Trp Thr Asp His Leu  
 485 490 495  
 His Leu Arg Ile Glu Arg Met His Trp Pro Gly Ser Gly Gln Gln Leu  
 500 505 510  
 Pro Arg Ser Ile Cys Ser Leu Pro Cys Gln Pro Gly Glu Arg Lys Lys  
 515 520 525  
 Thr Val Lys Gly Met Pro Cys Cys Trp His Cys Glu Pro Cys Thr Gly

530		535		540
Tyr Gln Tyr Gln Val Asp Arg Tyr Thr Cys Lys Thr Cys Pro Tyr Asp				
545		550		555
Met Arg Pro Thr Glu Asn Arg Thr Gly Cys Arg Pro Ile Pro Ile Ile				
		565		570
Lys Leu Glu Trp Gly Ser Pro Trp Ala Val Leu Pro Leu Phe Leu Ala				
		580		585
Val Val Gly Ile Ala Ala Thr Leu Phe Val Val Ile Thr Phe Val Arg				
		595		600
Tyr Asn Asp Thr Pro Ile Val Lys Ala Ser Gly Arg Glu Leu Ser Tyr				
		610		615
Val Leu Leu Ala Gly Ile Phe Leu Cys Tyr Ala Thr Thr Phe Leu Met				
625		630		635
Ile Ala Glu Pro Asp Leu Gly Thr Cys Ser Leu Arg Arg Ile Phe Leu				
		645		650
Gly Leu Gly Met Ser Ile Ser Tyr Ala Ala Leu Leu Thr Lys Thr Asn				
		660		665
Arg Ile Tyr Arg Ile Phe Glu Gln Gly Lys Arg Ser Val Ser Ala Pro				
		675		680
Arg Phe Ile Ser Pro Ala Ser Gln Leu Ala Ile Thr Phe Ser Leu Ile				
		690		695
Ser Leu Gln Leu Leu Gly Ile Cys Val Trp Phe Val Val Asp Pro Ser				
705		710		715
His Ser Val Val Asp Phe Gln Asp Gln Arg Thr Leu Asp Pro Arg Phe				
		725		730
Arg Val Leu Lys Cys Asp Ile Ser Asp Leu Ser Leu Ile Cys Leu Leu				
		740		745
Gly Tyr Ser Met Leu Leu Met Val Thr Cys Thr Val Tyr Ala Ile Lys				
		755		760
Thr Arg Gly Val Pro Glu Thr Phe Asn Glu Ala Lys Pro Ile Gly Phe				
		770		775
Thr Met Tyr Thr Thr Cys Ile Val Trp Leu Ala Phe Ile Pro Ile Phe				
785		790		795
Phe Gly Thr Ser Gln Ser Ala Asp Lys Leu Tyr Ile Gln Thr Thr Thr				
		805		810
Leu Thr Val Ser Val Ser Leu Ser Ala Ser Val Ser Leu Gly Met Leu				
		820		825
Tyr Met Pro Lys Val Tyr Ile Ile Leu Phe His Pro Glu Gln Asn Val				
		835		840
Pro Lys Arg Lys Arg Ser Leu Lys Ala Val Val Thr Ala Ala Thr Met				
		850		855
Ser Asn Lys Phe Thr Gln Lys Gly Asn Phe Arg Pro Asn Gly Glu Ala				
865		870		875
Lys Ser Glu Leu Cys Glu Asn Leu Glu Ala Pro Ala Thr Lys Gln Thr				
		885		890
Tyr Val Thr Tyr Thr Asn His Ala Ile				
		900		905

&lt;210&gt; 49

&lt;211&gt; 1416

&lt;212&gt; PRT

&lt;213&gt; Chimeric phmGluR4//CaR\*AAA\*G qi5

&lt;400&gt; 49

Met Val Cys Glu Gly Lys Arg Ser Ala Ser Cys Pro Cys Phe Phe Leu
1 5 10 15

Leu Thr Ala Lys Phe Tyr Trp Ile Leu Thr Met Met Gln Arg Thr His  
 20 25 30  
 Ser Gln Glu Tyr Ala His Ser Ile Arg Ile Asp Gly Asp Ile Thr Leu  
 35 40 45  
 Gly Gly Leu Phe Pro Val His Gly Arg Gly Ser Glu Gly Lys Pro Cys  
 50 55 60  
 Gly Glu Leu Lys Lys Glu Lys Gly Ile His Arg Leu Glu Ala Met Leu  
 65 70 75 80  
 Phe Ala Leu Asp Arg Ile Asn Asn Asp Pro Asp Leu Leu Pro Asn Ile  
 85 90 95  
 Thr Leu Gly Ala Arg Ile Leu Asp Thr Cys Ser Arg Asp Thr His Ala  
 100 105 110  
 Leu Glu Gln Ser Leu Thr Phe Val Gln Ala Leu Ile Glu Lys Asp Gly  
 115 120 125  
 Thr Glu Val Arg Cys Gly Ser Gly Gly Pro Pro Ile Ile Thr Lys Pro  
 130 135 140  
 Glu Arg Val Val Gly Val Ile Gly Ala Ser Gly Ser Ser Val Ser Ile  
 145 150 155 160  
 Met Val Ala Asn Ile Leu Arg Leu Phe Lys Ile Pro Gln Ile Ser Tyr  
 165 170 175  
 Ala Ser Thr Ala Pro Asp Leu Ser Asp Asn Ser Arg Tyr Asp Phe Phe  
 180 185 190  
 Ser Arg Val Val Pro Ser Asp Thr Tyr Gln Ala Gln Ala Met Val Asp  
 195 200 205  
 Ile Val Arg Ala Leu Lys Trp Asn Tyr Val Ser Thr Val Ala Ser Glu  
 210 215 220  
 Gly Ser Tyr Gly Glu Ser Gly Val Glu Ala Phe Ile Gln Lys Ser Arg  
 225 230 235 240  
 Glu Asp Gly Gly Val Cys Ile Ala Gln Ser Val Lys Ile Pro Arg Glu  
 245 250 255  
 Pro Lys Ala Gly Glu Phe Asp Lys Ile Ile Arg Arg Leu Leu Glu Thr  
 260 265 270  
 Ser Asn Ala Arg Ala Val Ile Ile Phe Ala Asn Glu Asp Asp Ile Arg  
 275 280 285  
 Arg Val Leu Glu Ala Ala Arg Arg Ala Asn Gln Thr Gly His Phe Phe  
 290 295 300  
 Trp Met Gly Ser Asp Ser Trp Gly Ser Lys Ile Ala Pro Val Leu His  
 305 310 315 320  
 Leu Glu Glu Val Ala Glu Gly Ala Val Thr Ile Leu Pro Lys Arg Met  
 325 330 335  
 Ser Val Arg Gly Phe Asp Arg Tyr Phe Ser Ser Arg Thr Leu Asp Asn  
 340 345 350  
 Asn Arg Arg Asn Ile Trp Phe Ala Glu Phe Trp Glu Asp Asn Phe His  
 355 360 365  
 Cys Lys Leu Ser Arg His Ala Leu Lys Lys Gly Ser His Val Lys Lys  
 370 375 380  
 Cys Thr Asn Arg Glu Arg Ile Gly Gln Asp Ser Ala Tyr Glu Gln Glu  
 385 390 395 400  
 Gly Lys Val Gln Phe Val Ile Asp Ala Val Tyr Ala Met Gly His Ala  
 405 410 415  
 Leu His Ala Met His Arg Asp Leu Cys Pro Gly Arg Val Gly Leu Cys  
 420 425 430  
 Pro Arg Met Asp Pro Val Asp Gly Thr Gln Leu Leu Lys Tyr Ile Arg  
 435 440 445  
 Asn Val Asn Phe Ser Gly Ile Ala Gly Asn Pro Val Thr Phe Asn Glu  
 450 455 460  
 Asn Gly Asp Ala Pro Gly Arg Tyr Asp Ile Tyr Gln Tyr Gln Leu Arg

465					470								480			
Asn	Asp	Ser	Ala	Glu	Tyr	Lys	Val	Ile	Gly	Ser	Trp	Thr	Asp	His	Leu	
				485					490					495		
His	Leu	Arg	Ile	Glu	Arg	Met	His	Trp	Pro	Gly	Ser	Gly	Gln	Gln	Leu	
				500					505					510		
Pro	Arg	Ser	Ile	Cys	Ser	Leu	Pro	Cys	Gln	Pro	Gly	Glu	Arg	Lys	Lys	
				515					520					525		
Thr	Val	Lys	Gly	Met	Pro	Cys	Cys	Trp	His	Cys	Glu	Pro	Cys	Thr	Gly	
				530					535					540		
Tyr	Gln	Tyr	Gln	Val	Asp	Arg	Tyr	Thr	Cys	Lys	Thr	Cys	Pro	Tyr	Asp	
545					550					555					560	
Met	Arg	Pro	Thr	Glu	Asn	Arg	Thr	Gly	Cys	Arg	Pro	Ile	Pro	Ile	Ile	
				565					570					575		
Lys	Leu	Glu	Trp	Gly	Ser	Pro	Trp	Ala	Val	Leu	Pro	Leu	Phe	Leu	Ala	
				580					585					590		
Val	Val	Gly	Ile	Ala	Ala	Thr	Leu	Phe	Val	Val	Ile	Thr	Phe	Val	Arg	
				595					600					605		
Tyr	Asn	Asp	Thr	Pro	Ile	Val	Lys	Ala	Ser	Gly	Arg	Glu	Leu	Ser	Tyr	
				610					615					620		
Val	Leu	Leu	Ala	Gly	Ile	Phe	Leu	Cys	Tyr	Ala	Thr	Thr	Phe	Leu	Met	
625					630					635					640	
Ile	Ala	Glu	Pro	Asp	Leu	Gly	Thr	Cys	Ser	Leu	Arg	Arg	Ile	Phe	Leu	
				645					650					655		
Gly	Leu	Gly	Met	Ser	Ile	Ser	Tyr	Ala	Ala	Leu	Leu	Thr	Lys	Thr	Asn	
				660					665					670		
Arg	Ile	Tyr	Arg	Ile	Phe	Glu	Gln	Gly	Lys	Arg	Ser	Val	Ser	Ala	Pro	
				675					680					685		
Arg	Phe	Ile	Ser	Pro	Ala	Ser	Gln	Leu	Ala	Ile	Thr	Phe	Ser	Leu	Ile	
				690					695					700		
Ser	Leu	Gln	Leu	Leu	Gly	Ile	Cys	Val	Trp	Phe	Val	Val	Asp	Pro	Ser	
705					710					715					720	
His	Ser	Val	Val	Asp	Phe	Gln	Asp	Gln	Arg	Thr	Leu	Asp	Pro	Arg	Phe	
				725					730					735		
Arg	Val	Leu	Lys	Cys	Asp	Ile	Ser	Asp	Leu	Ser	Leu	Ile	Cys	Leu	Leu	
				740					745					750		
Gly	Tyr	Ser	Met	Leu	Leu	Met	Val	Thr	Cys	Thr	Val	Tyr	Ala	Ile	Lys	
				755					760					765		
Thr	Arg	Gly	Val	Pro	Glu	Thr	Phe	Asn	Glu	Ala	Lys	Pro	Ile	Gly	Phe	
				770					775					780		
Thr	Met	Tyr	Thr	Thr	Cys	Ile	Val	Trp	Leu	Ala	Phe	Ile	Pro	Ile	Phe	
785					790					795					800	
Phe	Gly	Thr	Ser	Gln	Ser	Ala	Asp	Lys	Leu	Tyr	Ile	Gln	Thr	Thr	Thr	
				805					810					815		
Leu	Thr	Val	Ser	Val	Ser	Leu	Ser	Ala	Ser	Val	Ser	Leu	Gly	Met	Leu	
				820					825					830		
Tyr	Met	Pro	Lys	Val	Tyr	Ile	Ile	Leu	Phe	His	Pro	Glu	Gln	Asn	Thr	
				835					840					845		
Ile	Glu	Glu	Val	Arg	Cys	Ser	Thr	Ala	Ala	His	Ala	Phe	Lys	Val	Ala	
				850					855					860		
Ala	Arg	Ala	Thr	Leu	Arg	Arg	Ser	Asn	Val	Ser	Arg	Lys	Arg	Ser	Ser	
865					870					875					880	
Ser	Leu	Gly	Gly													

Thr Leu Pro Gln Gln Gln Arg Ser Gln Gln Gln Pro Arg Cys Lys Gln  
 930 935 940  
 Lys Val Ile Phe Gly Ser Gly Thr Val Thr Phe Ser Leu Ser Phe Asp  
 945 950 955 960  
 Glu Pro Gln Lys Asn Ala Met Ala His Gly Asn Ser Thr His Gln Asn  
 965 970 975  
 Ser Leu Glu Ala Gln Lys Ser Ser Asp Thr Leu Thr Arg His Gln Pro  
 980 985 990  
 Leu Leu Pro Leu Gln Cys Gly Glu Thr Asp Leu Asp Leu Thr Val Gln  
 995 1000 1005  
 Glu Thr Gly Leu Gln Gly Pro Val Gly Gly Asp Gln Arg Pro Glu Val  
 1010 1015 1020  
 Glu Asp Pro Glu Glu Leu Ser Pro Ala Leu Val Val Ser Ser Ser Gln  
 1025 1030 1035 1040  
 Ser Phe Val Ile Ser Gly Gly Gly Ser Thr Val Thr Glu Asn Val Val  
 1045 1050 1055  
 Asn Ser Ala Ala Ala Met Thr Leu Glu Ser Ile Met Ala Cys Cys Leu  
 1060 1065 1070  
 Ser Glu Glu Ala Lys Glu Ala Arg Arg Ile Asn Asp Glu Ile Glu Arg  
 1075 1080 1085  
 Gln Leu Arg Arg Asp Lys Arg Asp Ala Arg Arg Glu Leu Lys Leu Leu  
 1090 1095 1100  
 Leu Leu Gly Thr Gly Glu Ser Gly Lys Ser Thr Phe Ile Lys Gln Met  
 1105 1110 1115 1120  
 Arg Ile Ile His Gly Ser Gly Tyr Ser Asp Glu Asp Lys Arg Gly Phe  
 1125 1130 1135  
 Thr Lys Leu Val Tyr Gln Asn Ile Phe Thr Ala Met Gln Ala Met Ile  
 1140 1145 1150  
 Arg Ala Met Asp Thr Leu Lys Ile Pro Tyr Lys Tyr Glu His Asn Lys  
 1155 1160 1165  
 Ala His Ala Gln Leu Val Arg Glu Val Asp Val Glu Lys Val Ser Ala  
 1170 1175 1180  
 Phe Glu Asn Pro Tyr Val Asp Ala Ile Lys Ser Leu Trp Asn Asp Pro  
 1185 1190 1195 1200  
 Gly Ile Gln Glu Cys Tyr Asp Arg Arg Arg Glu Tyr Gln Leu Ser Asp  
 1205 1210 1215  
 Ser Thr Lys Tyr Tyr Leu Asn Asp Leu Asp Arg Val Ala Asp Pro Ala  
 1220 1225 1230  
 Tyr Leu Pro Thr Gln Gln Asp Val Leu Arg Val Arg Val Pro Thr Thr  
 1235 1240 1245  
 Gly Ile Ile Glu Tyr Pro Phe Asp Leu Gln Ser Val Ile Phe Arg Met  
 1250 1255 1260  
 Val Asp Val Gly Gly Gln Arg Ser Arg Lys Trp Ile His Cys Phe Glu  
 1265 1270 1275 1280  
 Asn Val Thr Ser Ile Met Phe Leu Val Ser Glu Tyr Asp Gln Val Leu  
 1285 1290 1295  
 Val Glu Ser Asp Asn Glu Asn Arg Met Glu Glu Ser Lys Ala Leu Phe  
 1300 1305 1310  
 Arg Thr Ile Ile Thr Tyr Pro Trp Phe Gln Asn Ser Ser Val Ile Leu  
 1315 1320 1325  
 Phe Leu Asn Lys Lys Asp Leu Leu Glu Glu Lys Ile Met Tyr Ser His  
 1330 1335 1340  
 Leu Val Asp Tyr Phe Pro Glu Tyr Asp Gly Pro Gln Arg Asp Ala Gln  
 1345 1350 1355 1360  
 Ala Ala Arg Glu Phe Ile Leu Lys Met Phe Val Asp Leu Asn Pro Asp  
 1365 1370 1375  
 Ser Asp Lys Ile Ile Tyr Ser His Phe Thr Cys Ala Thr Asp Thr Glu

	1380		1385		1390
Asn	Ile Arg Phe Val Phe Ala	Ala Val Lys Asp Thr	Ile Leu Gln Leu		
	1395	1400	1405		
Asn	Leu Lys Asp Cys Gly Leu Phe				
	1410	1415			

&lt;210&gt; 50

&lt;211&gt; 1411

&lt;212&gt; PRT

<213> Chimeric phmGluR8//CaR\*AAA\*G<sub>q</sub>i5

&lt;400&gt; 50

Met	Val	Cys	Glu	Gly	Lys	Arg	Ser	Ala	Ser	Cys	Pro	Cys	Phe	Phe	Leu
1				5					10					15	
Leu	Thr	Ala	Lys	Phe	Tyr	Trp	Ile	Leu	Thr	Met	Met	Gln	Arg	Thr	His
			20					25					30		
Ser	Gln	Glu	Tyr	Ala	His	Ser	Ile	Arg	Val	Asp	Gly	Asp	Ile	Ile	Leu
			35					40				45			
Gly	Gly	Leu	Phe	Pro	Val	His	Ala	Lys	Gly	Glu	Arg	Gly	Val	Pro	Cys
			50					55				60			
Gly	Glu	Leu	Lys	Lys	Glu	Lys	Gly	Ile	His	Arg	Leu	Glu	Ala	Met	Leu
65					70					75					80
Tyr	Ala	Ile	Asp	Gln	Ile	Asn	Lys	Asp	Pro	Asp	Leu	Leu	Ser	Asn	Ile
				85					90					95	
Thr	Leu	Gly	Val	Arg	Ile	Leu	Asp	Thr	Cys	Ser	Arg	Asp	Thr	Tyr	Ala
			100					105						110	
Leu	Glu	Gln	Ser	Leu	Thr	Phe	Val	Gln	Ala	Leu	Ile	Glu	Lys	Asp	Ala
			115					120					125		
Ser	Asp	Val	Lys	Cys	Ala	Asn	Gly	Asp	Pro	Pro	Ile	Phe	Thr	Lys	Pro
			130					135				140			
Asp	Lys	Ile	Ser	Gly	Val	Ile	Gly	Ala	Ala	Ala	Ser	Ser	Val	Ser	Ile
145					150					155					160
Met	Val	Ala	Asn	Ile	Leu	Arg	Leu	Phe	Lys	Ile	Pro	Gln	Ile	Ser	Tyr
				165					170					175	
Ala	Ser	Thr	Ala	Pro	Glu	Leu	Ser	Asp	Asn	Thr	Arg	Tyr	Asp	Phe	Phe
			180					185					190		
Ser	Arg	Val	Val	Pro	Pro	Asp	Ser	Tyr	Gln	Ala	Gln	Ala	Met	Val	Asp
			195				200					205			
Ile	Val	Thr	Ala	Leu	Gly	Trp	Asn	Tyr	Val	Ser	Thr	Leu	Ala	Ser	Glu
			210				215				220				
Gly	Asn	Tyr	Gly	Glu	Ser	Gly	Val	Glu	Ala	Phe	Thr	Gln	Ile	Ser	Arg
225					230					235					240
Glu	Ile	Gly	Gly	Val	Cys	Ile	Ala	Gln	Ser	Gln	Lys	Ile	Pro	Arg	Glu
				245					250					255	
Pro	Arg	Pro	Gly	Glu	Phe	Glu	Lys	Ile	Ile	Lys	Arg	Leu	Leu	Glu	Thr
			260					265					270		
Pro	Asn	Ala	Arg	Ala	Val	Ile	Met	Phe	Ala	Asn	Glu	Asp	Asp	Ile	Arg
			275					280				285			
Arg	Ile	Ala	Ala	Lys	Lys	Leu	Asn	Gln	Ser	Gly	His	Phe	Leu	Trp	Ile
			290				295				300				
Gly	Ser	Asp	Ser	Trp	Gly	Ser	Lys	Ile	Ala	Pro	Val	Tyr	Gln	Gln	Glu
305					310					315					320
Glu	Ile	Ala	Glu	Gly	Ala	Val	Thr	Ile	Leu	Pro	Lys	Arg	Ala	Ser	Ile
				325					330					335	
Asp	Gly	Phe	Asp	Arg	Tyr	Phe	Arg	Ser	Arg	Thr	Leu	Ala	Asn	Asn	Arg
			340					345					350		



Arg	Asn	Val	Trp	Phe	Ala	Glu	Phe	Trp	Glu	Glu	Asn	Phe	Gly	Cys	Lys
		355					360					365			
Leu	Gly	Ser	His	Gly	Lys	Arg	Asn	Ser	His	Ile	Lys	Lys	Cys	Thr	Gly
		370				375					380				
Leu	Glu	Arg	Ile	Ala	Arg	Asp	Ser	Ser	Tyr	Glu	Gln	Glu	Gly	Lys	Val
385						390					395				400
Gln	Phe	Val	Ile	Asp	Ala	Val	Tyr	Ser	Met	Ala	Tyr	Ala	Leu	His	Asn
				405					410					415	
Met	His	Lys	Asp	Leu	Cys	Pro	Gly	Tyr	Ile	Gly	Leu	Cys	Pro	Arg	Met
			420					425					430		
Ser	Thr	Ile	Asp	Gly	Lys	Glu	Leu	Leu	Gly	Tyr	Ile	Arg	Ala	Val	Asn
		435					440					445			
Phe	Asn	Gly	Ser	Ala	Gly	Thr	Pro	Val	Thr	Phe	Asn	Glu	Asn	Gly	Asp
	450					455					460				
Ala	Pro	Gly	Arg	Tyr	Asp	Ile	Phe	Gln	Tyr	Gln	Ile	Thr	Asn	Lys	Ser
465					470					475					480
Thr	Glu	Tyr	Lys	Val	Ile	Gly	His	Trp	Thr	Asn	Gln	Leu	His	Leu	Lys
				485					490					495	
Val	Glu	Asp	Met	Gln	Trp	Ala	His	Arg	Glu	His	Thr	His	Pro	Ala	Ser
			500					505					510		
Val	Cys	Ser	Leu	Pro	Cys	Lys	Pro	Gly	Glu	Arg	Lys	Lys	Thr	Val	Lys
		515					520					525			
Gly	Val	Pro	Cys	Cys	Trp	His	Cys	Glu	Arg	Cys	Glu	Gly	Tyr	Asn	Tyr
	530					535					540				
Gln	Val	Asp	Glu	Leu	Ser	Cys	Glu	Leu	Cys	Pro	Leu	Asp	Gln	Arg	Pro
545					550					555					560
Asn	Met	Asn	Arg	Thr	Gly	Cys	Gln	Leu	Ile	Pro	Ile	Ile	Lys	Leu	Glu
				565					570					575	
Trp	His	Ser	Pro	Trp	Ala	Val	Val	Pro	Val	Phe	Val	Ala	Ile	Leu	Gly
			580					585					590		
Ile	Ile	Ala	Thr	Thr	Phe	Val	Ile	Val	Thr	Phe	Val	Arg	Tyr	Asn	Asp
		595					600					605			
Thr	Pro	Ile	Val	Arg	Ala	Ser	Gly	Arg	Glu	Leu	Ser	Tyr	Val	Leu	Leu
	610					615					620				
Thr	Gly	Ile	Phe	Leu	Cys	Ile	Thr	Phe	Leu	Met	Ile	Ala	Ala	Pro	Asp
625					630					635					640
Thr	Ile	Ile	Cys	Ser	Phe	Arg	Arg	Val	Phe	Leu	Gly	Leu	Gly	Met	Cys
			645						650					655	
Phe	Ser	Tyr	Ala	Ala	Leu	Leu	Thr	Lys	Thr	Asn	Arg	Ile	His	Arg	Ile
			660					665					670		
Phe	Glu	Gln	Gly	Lys	Lys	Ser	Val	Thr	Ala	Pro	Lys	Phe	Ile	Ser	Pro
	675						680					685			
Ala	Ser	Gln	Leu	Val	Ile	Thr	Phe	Ser	Leu	Ile	Ser	Val	Gln	Leu	Leu
	690					695					700				
Gly	Val	Phe	Val	Trp	Phe	Val	Val	Asp	Pro	Pro	His	Ile	Ile	Ile	Asp
705					710					715					720
Tyr	Gly	Glu	Gln	Arg	Thr	Leu	Asp	Pro	Glu	Lys	Arg	Val	Leu	Lys	Cys
			725						730					735	
Asp	Ile	Ser	Asp	Leu	Ser	Leu	Ile	Cys	Ser	Leu	Gly	Tyr	Ser	Ile	Leu
			740					745					750		
Leu	Met	Val	Thr	Cys	Thr	Val	Tyr	Ala	Ile	Lys	Thr	Arg	Gly	Val	Pro
	755						760					765			
Glu	Thr	Phe	Asn	Glu	Ala	Lys	Pro	Ile	Gly	Phe	Thr	Met	Tyr	Thr	Thr
	770					775					780				
Cys	Ile	Ile	Trp	Leu	Ala	Phe	Ile	Pro	Ile	Phe	Phe	Gly	Thr	Ala	Gln
785					790					795					800
Ser	Ala	Glu	Lys	Met	Tyr	Ile	Gln	Thr	Thr	Thr	Leu	Thr	Val	Ser	Met

													805			810			815		
Ser	Leu	Ser	Ala	Ser	Val	Ser	Leu	Gly	Met	Leu	Tyr	Met	Pro	Lys	Val						
			820				825						830								
Tyr	Ile	Ile	Ile	Phe	His	Pro	Glu	Gln	Asn	Thr	Ile	Glu	Glu	Val	Arg						
			835				840						845								
Cys	Ser	Thr	Ala	Ala	His	Ala	Phe	Lys	Val	Ala	Ala	Arg	Ala	Thr	Leu						
			850				855						860								
Arg	Arg	Ser	Asn	Val	Ser	Arg	Lys	Arg	Ser	Ser	Ser	Leu	Gly	Gly	Ser						
865				870						875			880								
Thr	Gly	Ser	Thr	Pro	Ser	Ser	Ser	Ile	Ser	Ser	Lys	Ser	Asn	Ser	Glu						
			885						890			895									
Asp	Pro	Phe	Pro	Gln	Pro	Glu	Arg	Gln	Lys	Gln	Gln	Gln	Pro	Leu	Ala						
			900						905			910									
Leu	Thr	Gln	Gln	Glu	Gln	Gln	Gln	Gln	Pro	Leu	Thr	Leu	Pro	Gln	Gln						
			915						920			925									
Gln	Arg	Ser	Gln	Gln	Gln	Pro	Arg	Cys	Lys	Gln	Lys	Val	Ile	Phe	Gly						
			930						935			940									
Ser	Gly	Thr	Val	Thr	Phe	Ser	Leu	Ser	Phe	Asp	Glu	Pro	Gln	Lys	Asn						
945				950						955			960								
Ala	Met	Ala	His	Gly	Asn	Ser	Thr	His	Gln	Asn	Ser	Leu	Glu	Ala	Gln						
			965						970			975									
Lys	Ser	Ser	Asp	Thr	Leu	Thr	Arg	His	Gln	Pro	Leu	Leu	Pro	Leu	Gln						
			980						985			990									
Cys	Gly	Glu	Thr	Asp	Leu	Asp	Leu	Thr	Val	Gln	Glu	Thr	Gly	Leu	Gln						
			995						1000			1005									
Gly	Pro	Val	Gly	Gly	Asp	Gln	Arg	Pro	Glu	Val	Glu	Asp	Pro	Glu	Glu						
			1010						1015			1020									
Leu	Ser	Pro	Ala	Leu	Val	Val	Ser	Ser	Ser	Gln	Ser	Phe	Val	Ile	Ser						
1025				1030						1035			1040								
Gly	Gly	Gly	Ser	Thr	Val	Thr	Glu	Asn	Val	Val	Asn	Ser	Ala	Ala	Ala						
			1045						1050			1055									
Met	Thr	Leu	Glu	Ser	Ile	Met	Ala	Cys	Cys	Leu	Ser	Glu	Glu	Ala	Lys						
			1060						1065			1070									
Glu	Ala	Arg	Arg	Ile	Asn	Asp	Glu	Ile	Glu	Arg	Gln	Leu	Arg	Arg	Asp						
			1075						1080			1085									
Lys	Arg	Asp	Ala	Arg	Arg	Glu	Leu	Lys	Leu	Leu	Leu	Leu	Gly	Thr	Gly						
			1090						1095			1100									
Glu	Ser	Gly	Lys	Ser	Thr	Phe	Ile	Lys	Gln	Met	Arg	Ile	Ile	His	Gly						
1105				1110						1115			1120								
Ser	Gly	Tyr	Ser	Asp	Glu	Asp	Lys	Arg	Gly	Phe	Thr	Lys	Leu	Val	Tyr						
			1125						1130			1135									
Gln	Asn	Ile	Phe	Thr	Ala	Met	Gln	Ala	Met	Ile	Arg	Ala	Met	Asp	Thr						
			1140						1145			1150									
Leu	Lys	Ile	Pro	Tyr	Lys	Tyr	Glu	His	Asn	Lys	Ala	His	Ala	Gln	Leu						
			1155						1160			1165									
Val	Arg	Glu	Val	Asp	Val	Glu	Lys	Val	Ser	Ala	Phe	Glu	Asn	Pro	Tyr						
			1170						1175			1180									
Val	Asp	Ala	Ile	Lys	Ser	Leu	Trp	Asn	Asp	Pro	Gly	Ile	Gln	Glu	Cys						
1185				1190						1195			1200								
Tyr	Asp	Arg	Arg	Arg	Glu	Tyr	Gln														

Gln Arg Ser Arg Lys Trp Ile His Cys Phe Glu Asn Val Thr Ser Ile  
 1265 1270 1275 1280  
 Met Phe Leu Val Ser Glu Tyr Asp Gln Val Leu Val Glu Ser Asp Asn  
 1285 1290 1295  
 Glu Asn Arg Met Glu Glu Ser Lys Ala Leu Phe Arg Thr Ile Ile Thr  
 1300 1305 1310  
 Tyr Pro Trp Phe Gln Asn Ser Ser Val Ile Leu Phe Leu Asn Lys Lys  
 1315 1320 1325  
 Asp Leu Leu Glu Glu Lys Ile Met Tyr Ser His Leu Val Asp Tyr Phe  
 1330 1335 1340  
 Pro Glu Tyr Asp Gly Pro Gln Arg Asp Ala Gln Ala Ala Arg Glu Phe  
 1345 1350 1355 1360  
 Ile Leu Lys Met Phe Val Asp Leu Asn Pro Asp Ser Asp Lys Ile Ile  
 1365 1370 1375  
 Tyr Ser His Phe Thr Cys Ala Thr Asp Thr Glu Asn Ile Arg Phe Val  
 1380 1385 1390  
 Phe Ala Ala Val Lys Asp Thr Ile Leu Gln Leu Asn Leu Lys Asp Cys  
 1395 1400 1405  
 Gly Leu Phe  
 1410

1265 1270 1275 1280  
 1285 1290 1295  
 1300 1305 1310  
 1315 1320 1325  
 1330 1335 1340  
 1345 1350 1355 1360  
 1365 1370 1375  
 1380 1385 1390  
 1395 1400 1405  
 1410